## (19) World Intellectual Property Organization International Bureau



# 

(43) International Publication Date 7 October 2004 (07.10,2004)

**PCT** 

(10) International Publication Number WO 2004/085408 A1

- (51) International Patent Classification<sup>7</sup>: C07D 239/47, 401/14, 405/14, 409/14, 413/14, 417/14, 403/14, A61K 31/513, 31/5377, A61P 25/28
- (21) International Application Number:

PCT/JP2004/004320

- (22) International Filing Date: 26 March 2004 (26.03.2004)
- (25) Filing Language:

Fnolis

(26) Publication Language:

English

(30) Priority Data:

2003-126021

26 March 2003 (26.03.2003) JP

2003-126022 26 March 2003 (26.03.2003)

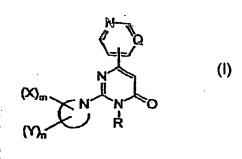
- (71) Applicants (for all designated States except US): MIT-SUBISHI PHARMA CORPORATION [JP/JP]; 6-9, Hiranomachi 2-chome, Chuo-ku, Osaka-shi, Osaka, 5410046 (JP). SANOFI-SYNTHELABO [FR/FR]; 174 AVENUE DE FRANCE PARIS, F-75013 (FR).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): WATANABE, Kazutoshi [JP/JP]; c/o MITSUBISHI PHARMA COR-PORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). UEHARA, Fumiaki [JP/JP]; c/o MITSUBISHI PHARMA COR-PORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). HIKI, Shinsuke [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION,

TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). YOKOSHIMA, Satoshi [JP/JP]: c/o MITSUBISHI PHARMA CORPORATION. TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). USUI, Yoshihiro [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). OKUYAMA, Masahiro [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OF-FICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). SHODA, Aya [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). ARITOMO, Keiichi [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). KOHARA, Toshiyuki [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP). FUKUNAGA, Kenji [JP/JP]; c/o MITSUBISHI PHARMA CORPORATION, TOKYO OFFICE, 2-6, Nihonbashi-honcho 2-chome, Chuo-ku, Tokyo, 1038405 (JP).

- (74) Agent: SIKS & CO., 8th Floor, Kyobashi-Nisshoku Bldg., 8-7, Kyobashi 1-chome, Chuo-ku, Tokyo, 1040031 (JP).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE,

[Continued on next page]

### (54) Title: 2, 3, 6-TRISUBSTITUTED-4-PYRIMIDONE DERIVATIVES



(57) Abstract: A pyrimidone derivative having tau protein kinase 1 inhibitory activity which is represented by formula (I) or a salt thereof, or a solvate thereof or a hydrate thereof; useful for prventive and/or therapeutic treatment of diseases such as neurodegenerative diseases (e.g. Alzheimer disease); wherein Q represents CH or nitrogen atom; R represents a  $C_1$ - $C_{12}$  alkyl group; the ring of Formula (I): represents piperazine ring or piperidine ring; each X independently represents a  $C_1$ - $C_8$  alkyl group, an optionally partially hydrogenated  $C_6$ - $C_{10}$  aryl ring, an indan ring or the like; m represents an integer of 1 to 3; each Y independently represents a halogen atom, a hydroxy group, a cyano group, a  $C_1$ - $C_6$  alkyl group or the like; n represents an integer of 0 to 8; when X and Y or two Y groups are attached on the same carbon atom, they may combine to each other to form a  $C_2$ - $C_6$  alkylene group.

**BEST AVAILABLE COPY** 

WO 2004/085408 A1

# 

KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

### Published:

with international search report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

### DESCRIPTION

#### 2,3,6-TRISUBSTITUTED -4-PYRIMIDONE DERIVATIVES

### Technical Field

The present invention relates to compounds that are useful as an active ingredient of a medicament for preventive and/or therapeutic treatment of diseases mainly caused by abnormal activity of tau protein kinase 1, such as neurodegenerative diseases (e.g. Alzheimer disease).

## **Background Art**

Alzheimer disease is progressive senile dementia, in which marked cerebral cortical atrophy is observed due to degeneration of nerve cells and decrease of nerve cell number. Pathologically, numerous senile plaques and neurofibrillary tangles are observed in brain. The number of patients has been increased with the increment of aged population, and the disease arises a serious social problem. Although various theories have been proposed, a cause of the disease has not yet been elucidated. Early resolution of the cause has been desired.

It has been known that the degree of appearance of two characteristic pathological changes of Alzheimer disease well correlates to the degree of intellectual dysfunction. Therefore, researches have been conducted from early 1980's to reveal the cause of the disease through molecular level investigations of components of the two pathological changes. Senile plaques accumulate extracellularly, and  $\beta$  amyloid protein has been elucidated as their main component (abbreviated as "A $\beta$ " hereinafter in the specification: Biochem. Biophys. Res. Commun., 120, 855 (1984); EMBO J., 4, 2757 (1985); Proc. Natl. Acad. Sci. USA, 82, 4245 (1985)). In the other pathological change, i.e., the neurofibrillary tangles, a double-helical filamentous substance called paired helical filament (abbreviated

as "PHF" hereinafter in the specification) accumulate intracellularly, and tau protein, which is a kind of microtubule-associated protein specific for brain, has been revealed as its main component (Proc. Natl. Acad. Sci. USA, 85, 4506 (1988); Neuron, 1, 827 (1988)).

Furthermore, on the basis of genetic investigations, presentlins 1 and 2 were found as causative genes of familial Alzheimer disease (Nature, 375, 754 (1995); Science, 269, 973 (1995); Nature. 376, 775 (1995)), and it has been revealed that presence of mutants of presentlins 1 and 2 promotes the secretion of A  $\beta$  (Neuron, 17, 1005 (1996); Proc. Natl. Acad. Sci. USA, 94, 2025 (1997)). From these results, it is considered that, in Alzheimer disease, A  $\beta$  abnormally accumulates and agglomerates due to a certain reason, which engages with the formation of PHF to cause death of nerve cells. It is also expected that extracellular outflow of glutamic acid and activation of glutamate receptor responding to the outflow may possibly be important factors in an early process of the nerve cell death caused by ischemic cerebrovascular accidents (Sai-shin Igaku [Latest Medicine], 49, 1506 (1994)).

It has been reported that kainic acid treatment that stimulates the AMPA receptor, one of glutamate receptor, increases mRNA of the amyloid precursor protein (abbreviated as "APP" hereinafter in the specification) as a precursor of A  $\beta$  (Society for Neuroscience Abstracts, 17, 1445 (1991)), and also promotes metabolism of APP (The Journal of Neuroscience, 10, 2400 (1990)). Therefore, it has been strongly suggested that the accumulation of A $\beta$  is involved in cellular death due to ischemic cerebrovascular disorders. Other diseases in which abnormal accumulation and agglomeration of A $\beta$  are observed include, for example, Down syndrome, cerebral bleeding due to solitary cerebral amyloid angiopathy, Lewy body disease (Shin-kei Shinpo [Nerve Advance], 34, 343 (1990); Tanpaku-shitu Kaku-san Koso [Protein, Nucleic Acid, Enzyme], 41, 1476 (1996)) and the like. Furthermore, as diseases showing neurofibrillary tangles due to the PHF accumulation, examples

include progressive supranuclear palsy, subacute sclerosing panencephalitic parkinsonism, postencephalitic parkinsonism, pugilistic encephalitis, Guam parkinsonism-dementia complex, Lewy body disease and the like (Tanpakushitu Kakusan Koso [Protein, Nucleic Acid, Enzyme], 36, 2 (1991); Igaku no Ayumi [Progress of Medicine], 158, 511 (1991); Tanpakushitu Kakusan Koso [Protein, Nucleic Acid, Enzyme], 41, 1476 (1996)).

The tau protein is generally composed of a group of related proteins that forms several bands at molecular weights of 48-65 kDa in SDS-polyacrylamide gel electrophoresis, and it promotes the formation of microtubules. It has been verified that tau protein incorporated in the PHF in the brain suffering from Alzheimer disease is abnormally phosphorylated compared with usual tau protein (J. Biochem., 99, 1807 (1986); Proc. Natl. Acad. Sci. USA, 83, 4913 (1986)). An enzyme catalyzing the abnormal phosphorylation has been isolated. The protein was named as tau protein kinase 1 (abbreviated as "TPK1" hereinafter in the specification), and its physicochemical properties have been elucidated (Seikagaku [Biochemistry], 64, 308 (1992); J. Biol. Chem., 267, 10897 (1992)). Moreover, cDNA of rat TPK1 was cloned from a rat cerebral cortex cDNA library based on a partial amino acid sequence of TPK1, and its nucleotide sequence was determined and an amino acid sequence was deduced (Japanese Patent Un-examined Publication [Kokai] No. 6-239893/1994). As a result, it has been revealed that the primary structure of the rat TPK1 corresponds to that of the enzyme known as rat GSK-3  $\beta$  (glycogen synthase kinase  $3\beta$ , FEBS Lett., 325, 167 (1993)).

It has been reported that A  $\beta$ , the main component of senile plaques, is neurotoxic (Science, 250, 279 (1990)). However, various theories have been proposed as for the reason why A  $\beta$  causes the cell death, and any authentic theory has not yet been established. Takashima et al. observed that the cell death was caused by A  $\beta$  treatment of fetal rat hippocampus primary culture system, and then found that the TPK1 activity was increased by A  $\beta$  treatment and the cell death by

A  $\beta$  was inhibited by antisense of TPK1 (Proc. Natl. Acad. Sci. USA, 90, 7789 (1993); Japanese Patent Un-examined Publication [Kokai] No. 6-329551/1994).

In view of the foregoing, compounds which inhibit the TPK1 activity may possibly suppress the neurotoxicity of A  $\beta$  and the formation of PHF and inhibit the nerve cell death in the Alzheimer disease, thereby cease or defer the progress of the disease. The compounds may also be possibly used as a medicament for therapeutic treatment of ischemic cerebrovascular disorder, Down syndrome, cerebral amyloid angiopathy, cerebral bleeding due to Lewy body disease and the like by suppressing the cytotoxicity of A  $\beta$ . Furthermore, the compounds may possibly be used as a medicament for therapeutic treatment of neurodegenerative diseases such as progressive supranuclear palsy, subacute sclerosing panencephalitic parkinsonism, postencephalitic parkinsonism, pugilistic encephalitis, Guam parkinsonism-dementia complex, Lewy body disease, Pick's disease, corticobasal degeneration, frontotemporal dementia, vascular dementia, acute stroke and traumatic injuries, brain and spinal cord trauma, peripheral neuropathies, retinopathies and glaucoma; non-insulin dependent diabetes (such as diabetes type II), and obesity, manic depressive illness, schizophrenia, alopecia, cancers such as breast cancer, non-small cell lung carcinoma, thyroid cancer, T or B-cell leukemia and several virus-induced tumors.

As structurally similar compounds to the compounds of the present invention represented by formula (I) described later, compounds represented by the following formula (A) are known:

wherein R represents 2,6-dichlorobenzyl group, 2-(2-chlorophenyl)ethylamino group, 3-phenylpropylamino group, or 1-methyl-3-phenylpropylamino group (WO98/24782). The compounds represented by formula (A) are characterized to have 4-fluorophenyl group at the 5-position of the pyrimidine ring and a hydroxy group at the 4-position, and not falling within the scope of the present invention. Moreover, main pharmacological activity of the compounds represented by formula (A) is anti-inflammatory effect, whereas the compounds of the present invention represented by formula (I) are useful as a TPK1 inhibitor or a medicament for therapeutic treatment of neurodegenerative diseases, and therefore, their pharmacological activities are totally different to each other.

Patent Document 1: WO 00/18758

· Patent Document 2: WO 01/70728

Patent Document 3: WO 01/70729

## Disclosure of the Invention

An object of the present invention is to provide compounds useful as an active ingredient of a medicament for preventive and/or therapeutic treatment of diseases such as Alzheimer disease. More specifically, the object is to provide novel compounds useful as an active ingredient of a medicament that enables radical prevention and/or treatment of the neurodegenerative diseases such as Alzheimer disease by inhibiting the TPK1 activity to suppress the neurotoxicity of A  $\beta$  and the formation of the PHF and by inhibiting the death of nerve cells.

In order to achieve the foregoing object, the inventors of the present invention conducted screenings of various compounds having inhibitory activity against the phosphorylation of TPK1. As a result, they found that compounds represented by the following formula (I) had the desired activity and were useful as an active ingredient of a medicament for preventive and/or therapeutic treatment of

WO 2004/085408

the aforementioned diseases. The present invention was achieved on the basis of these findings.

The present invention thus provides 3-substituted-4-pyrimidone derivatives represented by formula (I) or salts thereof, or solvates thereof or hydrates thereof:

$$(X)_{m}$$

$$(X)_{m}$$

$$(X)_{n}$$

$$(X)_$$

wherein Q represents CH or nitrogen atom;

R represents a C<sub>1</sub>-C<sub>12</sub> alkyl group which may be substituted; the ring of:

represents piperazine ring or piperidine ring; each X independently represents

 $X^1 - X^2 -$ 

wherein X¹ represents an oxo group; a C¹-C² alkyl group which may be substituted; a C³-C² cycloalkyl group which may be substituted; an optionally partially hydrogenated C²-C¹0 aryl ring which may be substituted; an indan ring which may be substituted; an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total; an aralkyloxy group; a group represented by -N(R²)(Rb) wherein R² and Rb are the same or different and each is hydrogen, a C¹-C² alkyl group which may be substituted, an aralkyl group which may be substituted, an aryl group which may be substituted, C¹-C² alkylcarbonyl group which may be

substituted,

C3-C8 cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted, C6-C10 arylcarbonyl group which may be substituted, C1-C8 alkysulfonyl group which may be substituted, C3-C8 cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted, C6-C10 arylsulfonyl group which may be substituted, C1-C8 alkyloxycarbonyl group which may be substituted, C3-C8 cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted, C6-C10 aryloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted, aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted,

N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N,N'-C8-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C<sub>8</sub>-C<sub>8</sub> cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted,

N,N'-diaralkylaminocarbonyl group which may be substituted,

N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

C6-C10 arylaminocarbonyl group which may be substituted,

N,N'-C6-C10 diarylaminocarbonyl group which may be substituted,

or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected

from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total; or Ra and Rb together with the adjacent nitrogen atom form a 4 to 7 membered heterocyclic ring which may further contain 1 to 4 groups selected from an oxygen atom, a sulfur atom, N-Rc (wherein Rc represents a hydrogen atom, a C1-C4 alkyl group which may be substituted, an aralkyl group which may be substituted, C3-C8 cycloalkyl group which may be substituted or an aryl group which may be substituted, C1-C8 alkylcarbonyl group which may be substituted, C3-C8 cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted, C6-C10 arylcarbonyl group which may be substituted, C1-C8 alkysulfonyl group which may be substituted, C3-C8 cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted, C<sub>6</sub>-C<sub>10</sub> arylsulfonyl group which may be substituted, C1-C8 alkyloxycarbonyl group which may be substituted, C3-C8 cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted, C6-C10 aryloxycarbonyl group which may be substituted, aminocarbonyl,

N-C<sub>1</sub>-C<sub>8</sub> alkylaminocarbonyl group which may be substituted, N, N'-C<sub>1</sub>-C<sub>8</sub> dialkylaminocarbonyl group which may be substituted,

 $N-C_1-C_8$  alkyl- $N'-C_3-C_8$  cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C<sub>3</sub>-C<sub>8</sub> cycloalkylaminocarbonyl group which may be substituted,

N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C<sub>8</sub>-C<sub>8</sub> cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted,
N,N'-diaralkylaminocarbonyl group which may be substituted,
N-aralkyl- N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted,
C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted,
N,N'-C<sub>6</sub>-C<sub>10</sub> diarylaminocarbonyl group which may be substituted,
or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total),
a carbonyl group, a sulfinyl group or a sulfonyl group in the ring, and said 4 to 7 membered heterocyclic ring may optionally be fused with an aryl group which may

be substituted;

X<sup>2</sup> represents a bond, a carbonyl group, a sulfinyl group, a sulfonyl group, an oxygen atom, a sulfur atom, a C1-C4 alkylene group which may be substituted or N-Rd (Rd represents a hydrogen atom, a C1-C4 alkyl group which may be substituted, an aralkyl group which may be substituted, Cs-Cs cycloalkyl group which may be substituted or an aryl group which may be substituted, C<sub>1</sub>-C<sub>8</sub> alkylcarbonyl group which may be substituted, C3-C8 cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted, C<sub>6</sub>-C<sub>10</sub> arylcarbonyl group which may be substituted, C<sub>1</sub>-C<sub>8</sub> alkysulfonyl group which may be substituted, C<sub>3</sub>-C<sub>8</sub> cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted, C6-C10 arylsulfonyl group which may be substituted, C1-C8 alkyloxycarbonyl group which may be substituted, C8-C8 cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted,

 $C_6$ - $C_{10}$  aryloxycarbonyl group which may be substituted, aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted,

N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C8-C8 cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C3-C6 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted,

aralkylaminocarbonyl group which may be substituted,

N,N'-diaralkylaminocarbonyl group which may be substituted,

N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

C6-C10 arylaminocarbonyl group which may be substituted,

N,N'-C6-C10 diarylaminocarbonyl group which may be substituted,

or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected

from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and

having 5 to 10 ring-constituting atoms in total);

m represents an integer of 1 to 3;

each Y independently represents a halogen atom, a hydroxy group, a cyano group, Y<sup>1</sup>-Y<sup>3</sup>- wherein Y<sup>1</sup> represents a C<sub>1</sub>-C<sub>8</sub> alkyl group which may be substituted; a C<sub>3</sub>-C<sub>8</sub> cycloalkyl group which may be substituted or a C<sub>6</sub>-C<sub>10</sub> aryl ring which may be substituted; Y<sup>3</sup> represents a carbonyl group, a sulfinyl group, a sulfonyl group, an oxygen atom, a sulfur atom, a C<sub>1</sub>-C<sub>4</sub> alkylene group which may be substituted or N-Re (Re represents a hydrogen atom, a C<sub>1</sub>-C<sub>4</sub> alkyl group which may be substituted, an aralkyl group which may be substituted, C<sub>3</sub>-C<sub>8</sub> cycloalkyl group which may be substituted or an aryl group which may be substituted,

C<sub>1</sub>-C<sub>8</sub> alkylcarbonyl group which may be substituted,
C<sub>3</sub>-C<sub>8</sub> cycloalkylcarbonyl group which may be substituted,
aralkycarbonyl group which may be substituted,
C<sub>6</sub>-C<sub>10</sub> arylcarbonyl group which may be substituted,
C<sub>1</sub>-C<sub>8</sub> alkysulfonyl group which may be substituted,
C<sub>3</sub>-C<sub>8</sub> cycloalkylsulfonyl group which may be substituted,
aralkysulfonyl group which may be substituted,
C<sub>6</sub>-C<sub>10</sub> arylsulfonyl group which may be substituted,
C<sub>1</sub>-C<sub>8</sub> alkyloxycarbonyl group which may be substituted,
C<sub>1</sub>-C<sub>8</sub> alkyloxycarbonyl group which may be substituted,
C<sub>3</sub>-C<sub>8</sub> cycloalkyloxycarbonyl group which may be substituted,
aralkyoxycarbonyl group which may be substituted,
C<sub>6</sub>-C<sub>10</sub> aryloxycarbonyl group which may be substituted,
aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted,

N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C3-C6 cycloalkylaminocarbonyl group which may be substituted,

N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C3-C8 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted,

N,N'-diaralkylaminocarbonyl group which may be substituted,

N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

C6-C10 arylaminocarbonyl group which may be substituted,

 $N,N'-C_6-C_{10}$  diarylaminocarbonyl group which may be substituted,

or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected

from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total),

n represents an integer of 0 to 8;

when X and Y or two Y groups are attached on the same carbon atom, they may combine to each other to form a  $C_2$ - $C_6$  alkylene group; and when m is 1, n is 0, and X is  $X^1$ -CO-,

- (1) X does not bind to 3-position of unsubstituted 1-piperazinyl group or does not bind to 3-position of a 4-alkyl-1-piperazinyl group; or
- (2) X does not bind to 3-position or 4-position of non-substituted 1-piperidinyl group.

According to another aspect of the present invention, there is provided a medicament comprising as an active ingredient a substance selected from the group consisting of the 3-substituted-4-pyrimidone derivatives represented by formula (I) and the physiologically acceptable salts thereof, and the solvates thereof and the hydrates thereof. As preferred embodiments of the medicament, there are provided the aforementioned medicament which is used for preventive and/or therapeutic treatment of diseases caused by tau protein kinase 1 hyperactivity, and the aforementioned medicament which is used for preventive and/or therapeutic treatment of neurodegenerative diseases.

As further preferred embodiments of the present invention, there are provided the aforementioned medicament wherein the diseases are selected from the group consisting of Alzheimer disease, ischemic cerebrovascular accidents, Down syndrome, cerebral bleeding due to cerebral amyloid angiopathy, progressive supranuclear palsy, subacute sclerosing panencephalitic parkinsonism, postencephalitic parkinsonism, pugilistic encephalitis, Guam parkinsonism-dementia complex, Lewy body disease, Pick's disease, corticobasal degeneration and frontotemporal dementia, vascular dementia, acute stroke and

traumatic injuries, brain and spinal cord trauma, peripheral neuropathies, retinopathies and glaucoma, non-insulin dependent diabetes (such as diabetes type II), and obesity, manic depressive illness, schizophrenia, alopecia, cancers such as breast cancer, non-small cell lung carcinoma, thyroid cancer, T or B-cell leukemia and several virus-induced tumors; and the aforementioned medicament in the form of pharmaceutical composition containing the above substance as an active ingredient together with one or more pharmaceutical additives.

The present invention further provides an inhibitor of tau protein kinase 1 comprising as an active ingredient a substance selected from the group consisting of the 3-substituted-4-pyrimidone derivatives of formula (I) and the salts thereof, and the solvates thereof and the hydrates thereof.

According to further aspects of the present invention, there are provided a method for preventive and/or therapeutic treatment of diseases caused by tau protein kinase 1 hyperactivity, which comprises the step of administering to a patient a preventively and/or therapeutically effective amount of a substance selected from the group consisting of the 3-substituted-4-pyrimidone derivatives of formula (I) and the physiologically acceptable salts thereof, and the solvates thereof and the hydrates thereof; and a use of a substance selected from the group consisting of the 3-substituted-4-pyrimidone derivatives of formula (I) and the physiologically acceptable salts thereof, and the solvates thereof and the hydrates thereof for the manufacture of the aforementioned medicament.

## Best Mode for Carrying Out the Invention

In the present specification, each group has the following meanings.

The alkyl group used herein may be either linear or branched.

The C<sub>1</sub>-C<sub>12</sub> alkyl group represented by R may be, for example, methyl group, ethyl group, n-propyl group, isopropyl group, n-butyl group, isobutyl group, sec-butyl group, tert-butyl group, n-pentyl group, isopentyl group, neopentyl group,

1,1-dimethylpropyl group, n-hexyl group, isohexyl group, or a linear or branched heptyl group, octyl group, nonyl group, decyl group, undecyl group or dodecyl group. Particularly preferred R is methyl group.

In the specification, when a functional group is defined as "which may be substituted" or "optionally substituted", the number of substituents as well as their types and substituting positions are not particularly limited, and when two or more substituents are present, they may be the same or different.

When the C<sub>1</sub>-C<sub>12</sub> alkyl group represented by R has one or more substituents, the alkyl group may have one or more substituents selected from, for example, the groups consisting of a C<sub>3</sub>-C<sub>8</sub> cycloalkyl group such as cyclopropyl group, cyclobutyl group, cyclopentyl group, cyclohexyl group, cycloheptyl group, cyclooctyl group; a C<sub>1</sub>-C<sub>5</sub> alkoxy group such as methoxy group, ethoxy group, propoxy group, isopropoxy group, butoxy group, isobutoxy group, tert-butoxy group; C<sub>1</sub>-C<sub>5</sub> alkylamino group or C<sub>2</sub>-C<sub>6</sub> dialkylamino group; a C<sub>6</sub>-C<sub>10</sub> aryl group such as phenyl group, 1-naphthyl group, and 2-naphthyl group.

The C<sub>1</sub>-C<sub>8</sub> alkyl group may be, for example, methyl group, ethyl group, n-propyl group, isopropyl group, n-butyl group, isobutyl group, sec-butyl group, tert-butyl group, n-pentyl group, isopentyl group, neopentyl group, 1,1-dimethylpropyl group, n-hexyl group, isohexyl group, or a linear or branched heptyl group or octyl group.

The C1-C4 alkyl group may be, for example, methyl group, ethyl group, n-propyl group, isopropyl group, n-butyl group, isobutyl group, sec-butyl group or tert-butyl group.

The  $C_8$ - $C_8$  cycloalkyl group may be, for example, cyclopropyl group, cyclobutyl group, cyclopentyl group, cyclohexyl group, cyclohexyl group or cycloctyl group.

The optionally partially hydrogenated  $C_6$ - $C_{10}$  aryl ring may be, for example a benzene ring, a naphthalene ring, an indan ring or a

PCT/JP2004/004320

# 1,2,3,4-tetrahydronaphthalene ring.

The heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total may be, for example, furan ring, dihydrofuran ring, tetrahydrofuran ring, pyran ring, dihydropyran ring, tetrahydropyran ring, benzofuran ring, dihydrobenzofuran, isobenzofuran ring, benzodioxol ring, chromene ring, chroman ring, isochroman ring, thiophene ring, benzothiophene ring, pyrrole ring, pyrroline ring, pyrrolidine ring, 2-oxopyrrolidine ring, imidazole ring, imidazoline ring, imidazolidine ring, pyrazole ring, pyrazoline ring, pyrazolidine ring, triazole ring, tetrazole ring, pyridine ring, pyridine oxide ring, piperidine ring, 4-oxopiperidine ring, pyrazine ring, piperazine ring, homopiperazine ring, pyrimidine ring, pyridazine ring, indole ring, indoline ring, isoindole ring, isoindoline ring, indazole ring, benzimidazole ring, benzotriazole ring, tetrahydroisoquinoline ring, benzothiazolinone ring, benzoxazolinone ring, purine ring, quinolizine ring, quinoline ring, phthalazine ring, naphthyridine ring, quinoxaline ring, quinazoline ring, cinnoline ring, pteridine ring, oxazole ring, oxazolidine ring, isoxazole ring, isoxazolidine ring, oxadiazole ring, thiazole ring, benzothiazole ring, thiazylidine ring, isothiazole ring, isothiazolidine ring, benzodioxole ring, dioxane ring, benzodioxane ring, dithian ring, morpholine ring, thiomorpholine ring, or phthalimide ring.

The aralkyl group may be, for example, benzyl group, 2-phenylethyl group, 3-phenylpropyl group or 4-phenylbutyl group.

The  $C_1$ - $C_4$  alkylene group may be, for example, methylene, ethylene, trimethylene or tetramethylene.

The 4 to 7 membered heterocyclic ring which may further contain 1 to 4 groups may be, for example, pyrrolidine, piperidine, morpholine, thiomorpholine, piperazine, homopiperazine, 2-oxopyrrolidine, pyrrole, imidazoline, imidazole, pyrazole, pyrroline, pyrrolidine, imidazolidine, imidazolone, succinimide or

glutarimide.

The  $C_6$ - $C_{10}$  aryl ring may be, for example, a benzene ring or a naphthalene ring, and the aryl group or the  $C_6$ - $C_{10}$  aryl group may be, for example, a phenyl group or naphthyl group.

When the ring represented by X or X1 has one or more substituents, the ring may have one or more substituents selected from the group consisting of a C1-C5 alkyl group such as methyl group, ethyl group, propyl group, isopropyl group, butyl group, isobutyl group, sec-butyl group, tert-butyl group, pentyl group, isopentyl group, neopentyl group, 1,1-dimethylpropyl group; C3-C6 cycloalkyl group such as cyclopropyl group, cyclobutyl group, cyclopentyl group, cyclohexyl group; a C3-C6 cycloalkyl-C1-C4 alkyl group such as cyclopropylmethyl, cyclopentylmethyl, cyclohexylmethyl; a C1-C4 hydroxyalkyl group such as hydroxymethyl, hydroxyethyl, hydroxypropyl; a halogen atom such as fluorine atom, chlorine atom, bromine atom, and iodine atom; a C1-C5 halogenated alkyl group such as trifluoromethyl group; hydroxyl group; cyano group; nitro group; formyl group; a benzene ring which may be substituted; a naphthalene ring which may be substituted; an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom and nitrogen atom, and having 5 to 10 ring-constituting atoms in total (same as the above); an amino group; an N- C3-C6 cycloalkyl-N-C1-C4 alkylaminoalkyl group wherein said C1-C4 alkyl may be substituted by hydroxy group or C1-C4 alkoxy group such as N-cyclopropyl-N-methylaminomethyl group, N-cyclohexyl-N-methylaminomethyl group; a C<sub>1</sub>-C<sub>5</sub> monoalkylaminomethyl group such as methylaminomethyl group, ethylaminomethyl group, propylaminomethyl group, isoproylaminomethyl group, butylaminomethyl group, isobutylaminomethyl group, tert-butylaminomethyl group, pentylaminomethyl group, isopentylaminomethyl group; a C2-C10 dialkylaminomethyl group such as dimethylaminomethyl group, diethylaminomethyl group, ethylmethylaminomethyl group,

methylpropylaminomethyl group; pyrrolidinylmethyl group; piperidinylmethyl group; morpholinomethyl group; piperazinylmethyl group; pyrrolylmethyl group; imidazolylmethyl group; pyrazolylmethyl group; triazolylmethyl group; and a group of the formula -E-Rf wherein E represents O, S, SO, SO2, CO or N(R4) and Rf represents a  $C_1$ - $C_5$  alkyl group (same as the above), a  $C_4$ - $C_7$  cycloalkyl group (same as the above), a C4-C7 cycloalkylalkl group (same as the above), a C1-C5 hydroxyalkyl group (same as the above), a benzene ring which may be substituted, a naphthalene ring which may be substituted, an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom and nitrogen atom, and having 5 to 10 ring-constituting atoms in total (same as the above), an N-C<sub>3</sub>-C<sub>6</sub> cycloalkyl-N-C<sub>1</sub>-C<sub>4</sub> alkylaminoalkyl group (same as the above), a  $C_1$ - $C_5$  monoalkylaminoalkyl group (same as the above), C2-C10 dialkylaminoalkyl group (same as the above), pyrrolidinylmethyl group, piperidinylmethyl group, morpholinomethyl group, piperazinylmethyl group, pyrrolylmethyl group, imidazolylmethyl group, pyrazolylmethyl group or triazolylmethyl group,

C1-C8 alkylcarbonyl group which may be substituted,
C3-C8 cycloalkylcarbonyl group which may be substituted,
aralkycarbonyl group which may be substituted,
C6-C10 arylcarbonyl group which may be substituted,
C1-C8 alkysulfonyl group which may be substituted,
C3-C8 cycloalkylsulfonyl group which may be substituted,
aralkysulfonyl group which may be substituted,
C6-C10 arylsulfonyl group which may be substituted,
C1-C8 alkyloxycarbonyl group which may be substituted,
C1-C8 cycloalkyloxycarbonyl group which may be substituted,
C3-C8 cycloalkyloxycarbonyl group which may be substituted,
aralkyoxycarbonyl group which may be substituted,

aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted,

N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C<sub>3</sub>-C<sub>6</sub> cycloalkylaminocarbonyl group which may be substituted,

N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted,

aralkylaminocarbonyl group which may be substituted,

N,N'-diaralkylaminocarbonyl group which may be substituted,

N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

C6-C10 arylaminocarbonyl group which may be substituted,

N,N'-C6-C10 diarylaminocarbonyl group which may be substituted,

and R4 represents a hydrogen atom, a C1-C4 alkyl group which may be substituted,

an aralkyl group which may be substituted, C3-C8 cycloalkyl group which may be

substituted or an aryl group which may be substituted,

C<sub>1</sub>-C<sub>8</sub> alkylcarbonyl group which may be substituted, .

C<sub>8</sub>-C<sub>8</sub> cycloalkylcarbonyl group which may be substituted,

aralkycarbonyl group which may be substituted,

C6-C10 arylcarbonyl group which may be substituted,

C1-C8 alkysulfonyl group which may be substituted,

C<sub>8</sub>-C<sub>8</sub> cycloalkylsulfonyl group which may be substituted,

aralkysulfonyl group which may be substituted,

C6-C10 arylsulfonyl group which may be substituted,

C1-C8 alkyloxycarbonyl group which may be substituted,

C3-C8 cycloalkyloxycarbonyl group which may be substituted,

aralkyoxycarbonyl group which may be substituted, C6-C10 aryloxycarbonyl group which may be substituted, aminocarbonyl,

having 5 to 10 ring-constituting atoms in total.

aminocarbonyl, N-C1-C8 alkylaminocarbonyl group which may be substituted, N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted, N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted, N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted, N-C1-C3 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted, C3-C8 cycloalkylaminocarbonyl group which may be substituted, N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted, N-C3-C8 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted, N-C3-C8 cycloalkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted, N,N'-diaralkylaminocarbonyl group which may be substituted, N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted, C6-C10 arylaminocarbonyl group which may be substituted, N,N'-C6-C10 diarylaminocarbonyl group which may be substituted, or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and

When the C<sub>6</sub>-C<sub>10</sub> aryl ring represented by Y<sup>1</sup> has one or more substituents, the ring may be substituted by one or more substituents selected from the groups consisting of halogen atoms, a C<sub>1</sub>-C<sub>5</sub> alkyl group, a C<sub>3</sub>-C<sub>6</sub> cycloalkyl group, a C<sub>3</sub>-C<sub>6</sub> cycloalkyloxy group, a C<sub>1</sub>-C<sub>5</sub> alkoxy group, a C<sub>4</sub>-C<sub>7</sub> cycloalkylalkoxy, a C<sub>1</sub>-C<sub>5</sub> alkylthio group, a C<sub>1</sub>-C<sub>5</sub> alkylsulfonyl group, a C<sub>1</sub>-C<sub>5</sub> halogenated alkyl, and a benzene ring.

When the ring represented by X, X<sup>1</sup> or Y<sup>1</sup> has one or more substituents, the substituent may further have one or more substituents selected from the group

consisting of a C1-C5 alkyl group such as methyl group, ethyl group, propyl group, isopropyl group, butyl group, isobutyl group, sec-butyl group, tert-butyl group, pentyl group, isopentyl group, neopentyl group, 1,1-dimethylpropyl group; C3-C6 cycloalkyl group such as cyclopropyl group, cyclobutyl group, cyclopentyl group, cyclohexyl group; a C8-C6 cycloalkyloxy group such as cyclopropyloxy group, cyclobutyloxy group, cyclopentyloxy group, cyclohexyloxy group; C1-C4 hydroxyalkyl group such as hydroxymethyl group, hydroxyethyl group, hydroxypropyl group, hydroxybutyl group; a C1-C5 alkoxy group such as methoxy group, ethoxy group, propoxy group, isopropoxy group, butoxy group, isobutoxy group, tert-butoxy group, pentyloxy group, and isopentyloxy group; a C4-C7 cycloalkylalkoxy group such as cyclopropylmethoxy group, cyclopentylmethoxy group; a C1-C5 alkylthio group such as methylthio group, ethylthio group, propylthio group, butylthio group, and pentylthio group; a C1-C5 alkylsulfonyl group such as methanesulfonyl group, ethanesulfonyl group, propanesulfonyl group, butanesulfonyl group, and pentanesulfonyl group; a halogen atom such as fluorine atom, chlorine atom, bromine atom, and iodine atom; a C1-C5 halogenated alkyl group such as trifluoromethyl group; a C1-C5 halogenated alkoxy group such as trifluoromethoxy group, 2,2,2-trifluoroethoxy group; hydroxyl group; cyano group; nitro group; formyl group; a C2-C6 alkylcarbonyl group such as acetyl group, propionyl group, butyryl group, and valeryl group; amino group; a C1-C5 monoalkylamino group such as methylamino group, ethylamino group, propylamino group, isopropylamino group, butylamino group, isobutylamino group, tert-butylamino group, pentylamino group, and isopentylamino group; a C2-C10 dialkylamino group such as dimethylamino group, ethylmethylamino group, diethylamino group, methylpropylamino group, and disopropylamino group; a cyclic amino group such as pyrrolidinyl group, piperidino group, morpholino group; a C2-C10 monoalkylaminomethyl group such as methylaminomethyl group, ethylaminomethyl group, propylaminomethyl group, isoproylaminomethyl group, butylaminomethyl group, isobutylaminomethyl group,

tert-butylaminomethyl group, pentylaminomethyl group, isopentylaminomethyl; a C<sub>3</sub>-C<sub>11</sub> dialkylaminomethyl group such as dimethylaminomethyl group, diethylaminomethyl group, ethylmethylaminomethyl group, methylpropylaminomethyl group; a phenyl group; an aralkyloxy group such as benzyloxy, 2-phenylethyloxy, 3-phenylpropyloxy; an aralkyloxycarbonyl group such as benzyloxycarbonyl, 2-phenylehoxycarbonyl; an C2-C4 alkanoyloxy-C1-C4 alkyl group such as acetyloxymethyl, 2-acetyloxyethyl, 2-propionyloxyethyl; an alkanoylamino group such as acetylamino, propionylamino, butyrylamino; N-C1-C4 alkyl-N-alkanoylamino group such as N-methyl-N-acetylamino, N-ethyl-N-acetylamino, N-methyl-N-propionylamino, N-methyl-N-butyrylamino; a heterocyclic ring amino group such as pyridylamino, pyrimidinylamino, thienylamino, furylamino; N-C1-C4 alkyl-N-heterocyclic ring amino group such as N-methyl-N-pyridylamino, N-methyl-N-pyrimidinylamino, N-methyl-N-thienylamino, N-methyl-N-furylamino; a diheterocyclic ring amino group such as dipyridylamino, dipyrimidinylamino, dithienylamino, difurylamino, and the like.

R may preferably be a  $C_1$ - $C_3$  alkyl group, more preferably a methyl group or an ethyl group. The substituent of the alkyl group may preferably be a  $C_3$ - $C_8$  alkyl group.

X may preferably be a benzene ring which may be substituted, a benzyl group which may be substituted, a naphthyl group which may be substituted, a benzofuran ring which may be substituted, a dihydrobenzofuran ring which may be substituted, a benzisoxazole ring which may be substituted, a benzisoxazole ring which may be substituted, a benzisothiazole ring which may be substituted, a benzisothiazole ring which may be substituted, and a benzopyrazole ring which may be substituted; more preferably a benzene ring which may be substituted, a benzyl group which may be substituted. Substituent of X may preferably be selected from the group consisting of a halogen

atom, a C1-C4 alkyl group, a C1-C4 alkoxy group, a hydroxy group, a nitro group, a cyano group, a perhalogenated C1-C4 alkyl group, a carboxyl group, a C1-C4 alkoxycarbonyl group, a C1-C4 alkylthio group, a C1-C4 alkoxysulfonyl group, amino group which may be substituted by a C1-C4 alkyl group, a benzene ring which may be substituted, and a cyclic amino group which may be substituted.

The compounds represented by the aforementioned formula (I) may form a salt. Examples of the salt include, when an acidic group exists, salts of alkali metals and alkaline earth metals such as lithium, sodium, potassium, magnesium, and calcium; salts of ammonia and amines such as methylamine, dimethylamine, trimethylamine, dicyclohexylamine, tris(hydroxymethyl)aminomethane,

N,N-bis(hydroxyethyl)piperazine, 2-amino-2-methyl-1-propanol, ethanolamine,

N-methylglucamine, and L-glucamine; or salts with basic amino acids such as lysine, δ-hydroxylysine, and arginine. When a basic group exists, examples include salts with mineral acids such as hydrochloric acid, hydrobromic acid, sulfuric acid, nitric acid, phosphoric acid; salts with organic acids such as methanesulfonic acid, benzenesulfonic acid, p-toluenesulfonic acid, acetic acid, propionic acid, tartaric acid, fumaric acid, maleic acid, malic acid, oxalic acid, succinic acid, citric acid, benzoic acid, mandelic acid, cinnamic acid, lactic acid, glycolic acid, glucuronic acid, ascorbic acid, nicotinic acid, and salicylic acid; or salts with acidic amino acids such as aspartic acid, and glutamic acid.

In addition to the 3-substituted-4-pyrimidone derivatives represented by the aforementioned formula (I) and salts thereof, their solvates and hydrates also fall within the scope of the present invention. The 3-substituted-4-pyrimidone derivatives represented by the aforementioned formula (I) may have one or more asymmetric carbon atoms. As for the stereochemistry of such asymmetric carbon atoms, they may independently be in either (R) and (S) configuration, and the pyrimidone derivative may exist as stereoisomers such as optical isomers, or diastereoisomers. Any stereoisomers in a pure form, any mixtures of stereoisomers,

racemates and the like fall within the scope of the present invention.

Preferred compounds of the present invention are represented by formula (II):

$$(X)_{p} \qquad (II)$$

$$(X)_{q} \qquad (Y)_{r}$$

wherein Q, R, X, Y are the same as those defined above; p is 0 or 1; q is 0 or 1; r is an integer of 0 to 6; p+q is 1 or 2;

and Z represents N or CZ1 wherein Z1 represents hydrogen atom or Y.

Examples of more preferred classes of compounds represented by formula (II) include:

- those wherein R represents a C<sub>1</sub>-C<sub>3</sub> alkyl group which may be substituted by a
   C<sub>3</sub>-C<sub>8</sub> cycloalkyl group;
- (2) the compounds of the above (1) wherein R is methyl group or ethyl group; Y is in 3-, 4- or 5-position of the piperazine ring or the piperidine ring; p+q is 1; and r is an integer of 0 to 3;
- (3) the compounds of the above (2) wherein X is a C<sub>1</sub>-C<sub>8</sub> alkyl group which may be substituted or a C<sub>6</sub>-C<sub>10</sub> aryl ring which may be substituted; Y is a C<sub>1</sub>-C<sub>6</sub> alkyl group which may be substituted; p is 1; q is 0; r is an integer of 0 to 3; and Z is N or CH; (4) the compounds of the above (3) wherein X is a benzene ring which may be substituted, a benzyl group which may be substituted; Y is a methyl group which may be substituted; Z is N and r is 0 or 1;
- (5) the compounds of the above (2) wherein X is a benzene ring which may be substituted, a benzyl group which may be substituted, a benzoyl group which may be substituted, or a benzisothiazol ring which may be substituted; Y is a methyl

group which may be substituted; Z is N and p is 0;

(6) the compounds of the above (2) wherein X is a  $C_1$ - $C_8$  alkyl group substituted by a benzene ring which may be substituted or a benzene ring which may be substituted; Y is a hydroxy group, a cyano group, or  $Y^1$ -CO- wherein  $Y^1$  is a  $C_1$ - $C_8$  alkyl group; Z is CH or C-Y and r is 0 or 1; and

(7) the compounds of the above (6) wherein X is a benzyl group which may be substituted or a benzene ring which may be substituted; Y is a hydroxy group, a cyano group, or an acetyl group; Z is CH or C-Y and r is 0 or 1.

Examples of particularly preferred classes of compounds represented by formula (II) include:

- (1) those wherein R is methyl group, Y is CH<sub>3</sub>O-CO- group or CH<sub>3</sub>CH<sub>2</sub>O-CO- group, Z is N, p is 0, q is 1, r is 0 or 1 and Y is in 3-position of the piperazine ring;
- (2) those wherein R is methyl group, Y is methyl group, benzyl group or acetyl group, Z is N, p is 1, q is 0, r is 0 or 1 and Y is in 4-position of the piperazine ring;
- (3) those wherein R is methyl group, Y is methyl group, Z is N, p is 1, q is 0, r is 1 to 3 and Y is in 3-, 4-, or 5-position of the piperazine ring;
- (4) those wherein R is methyl group, Y is hydroxyl group or cyano group, Z is CH, p is 1, q is 0, r is 0 or 1 and X and Y are attached on the same carbon atom;
- (5) those wherein R is methyl group, Y is hydroxyl group, cyano group or acetyl group, Z is C-Y, p is 0, q is 1 and r is 1.

Examples of preferred compounds of the present invention are shown in the tables below. However, the scope of the present invention is not limited to the following compounds.

Table-1		·				
	R1	R2	R3	R4	R5	R6
No.	CH3-	H	H	CH3-	H	H
XA2	CH3-	н	Н	СНЗСН2-	Н	Н
ХАЗ	снз-	Н	н	<b>^</b> \	Н	H-
XA4	снз-	н	н	<u> </u>	н	н
XA5	снз-	н	н	<b>√</b>	н	Н
XA6	снз-	н	н	人、	Н	н
XA7	снз–	Н	н		н	н
XA8	снз-	Н	н	<b>Y</b>	н	н
XA9	снз-	н	н	^\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н.
XA10	снз-	н	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA11	снз-	н	н	<u> </u>	н	н
XA12	снз-	н	н	<u> </u>	н .	н .
XA13	снз-	н	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA14	снз-	н	H.	<u></u>	н	н
XA15	снз-	Н	н	~~~`\	н	Н
XA16	снз-	н	н	Y~~~	н	н
XA17	снз-	н	н	n-C8H17-	Н	н
XA18	CH3-	н	н	<u> </u>	н .	Н
XA19	снз-	н	н	Qu	н	н
XA20	снз-	н	н		н	н
XA21	снэ–	н	н	Q	н .	н
XA22	снз-	н	н	<b>▷</b>	н	Н
XA23	снэ-	н	н	$\Diamond$	Н	н
XA24	снз-	н	Н	$\bigcirc$	н	н
XA25	снз-	н	н	$\bigcirc \dashv$	н	н



No.	R1	R2	R3	R4	R5	R6
XA26	снз-	н	Н	$\bigcirc$ $\vdash$	н	н
XA27	СН3-	н	Н		н	н
XA28	СН3-	н	н		н	н
XA29	СН3-	н	н		н	н
XA30	снз-	н	н	F-()-1	н	H
XA31	снз-	Н	н	CI	н	н
XA32	снз-	н	н	CI.	н	H ·
XA33	СН3-	Н	н	c <del></del>	н	Н
· XA34	СН3	Н	н	Br {.	н	н
XA35	снз-	н	н	Br.	н	н
XA36	СН3-	H	Н	Br-C>{	н .	н
XA37	СН3-	н	н		Н	Н
XA38	СН3-	н	н		н	Н
XA39	снз-	Н	н		н	Н
XA40	снз-	н	н	CH₃ С⊢₁	н	н
XA41	снз-	н	н	H <sub>3</sub> C	н	H
XA42	снз-	H	н	H3C-{}{	н	Н
XA43 ,	снз-	н	н		н	н
XA44	СН3-	н	Н	n-C <sub>3</sub> H <sub>7</sub> -{_}	н	н
XA45	снз-	н	н	n-C <sub>4</sub> H <sub>9</sub> -	н	н
XA46	снз	н	н	ОН	н	н
XA47	снз-	Н	н	HO —	н	н

No.	RI	R2	R3	R4	R5	R6
XA48	CH3-	Н	Н	HO-{\(\){\}	н	н
XA49	CH3-	н	н	OCH₃	н	н
XA50	снз-	н	н	H₃CO ⟨_}_	н	н
XA51	CH3-	н	Н	H₃CO- <b>⟨</b> }-{	н	н
XA52	CH3-	н	Н	C <sub>2</sub> H <sub>5</sub> O-{}-{	Н	н
XA53	CH3-	н	Н	n-C <sub>3</sub> H <sub>7</sub> O-{_}-{	Н	Н
XA54	CH3-	н	н	n-C <sub>4</sub> H <sub>9</sub> O-	Н	н
XA55	CH3-	н	н	NO <sub>2</sub>	Н	н
XA56	CH3-	н	н	O₂N △	н	Н
XA57	снз-	н	Н	O <sub>2</sub> N-{}	н	н
XA58	снз-	Н	Н	CN	н	н
XA59	снз-	н	н	NC	Н	н
XA60	снз-	Н	Н	NC-{\rightarrow}-{\rightarrow}-{\rightarrow}	Н	н
XA61	снз-	н .	Н	CF <sub>3</sub>	Н	н
XA62	снз-	н	н	F <sub>3</sub> C	н	н
XA63	снз-	H.	н	F <sub>3</sub> C-{}-{	н	н
XA64	CH3-	н	н	ÇOOH	Н	н
XA65	снз-	Н	н	HOOC.	Н	Н
XA66	снз-	Н	н	HOOC-{_}-	Н	Н
XA67	снз–	Н	н	CO <sub>2</sub> Me	н	н
XA68	снз-	H	н	MeO <sub>2</sub> C	н	н
XA69	снз-	Н	н	MeO <sub>2</sub> C-{{}	н	н



No.	R1	R2	R3	R4	R5	R6
XA70	снз–	н	н	CO <sub>2</sub> Et	н	н
XA71	CH3-	н	н	EtO <sub>2</sub> C	н	н
XA72	снз-	Н	н	EtO <sub>2</sub> C-	н	н
XA73	снз-	н	н	SMe	н	н
XA74	CH3-	Н	н	MeS	Н	H.
XA75	снз-	н	н	MeS-()-(	н	н
XA76	снз-	н	н	SO₂Me	Н	Н
XA77	CH3-	Н	н	MeO <sub>2</sub> S	н	н
XA78	снз-	н	н	MeO <sub>2</sub> S-{}-{	н	н
XA79	СН3-	н	н	NH <sub>2</sub>	Н	н
XA80	снз-	н .	Н	H <sub>2</sub> N	н	Н
XA81	снз-	н .	н	H <sub>2</sub> N-{}-{	н	н
XA82	снз-	Н	н	NMe <sub>2</sub>	Н	н
XA83	СН3-	н	н	Me <sub>2</sub> N .	н	н
XA84	снз-	н	н	Me <sub>2</sub> N-{¯}-{	н	н
XA85	снз-	н	н		н	н
XA86	снз-	н	н	CCT'	н	н
XA87	CH3-	Н	н	H H H	н	н
XA88	снз-	Н	н.		н	н
XA89	снз-	н	н	(D)	н	н
XA90	снз-	н	н	67	н	н
XA91	снз-	н	Н	S. i	н	н

No.	R1	R2	R3	R4	R5	R6
XA92	СН3-	н	н	\$\$\frac{1}{2}	н	н
XA93	СН3-	н	н	HNN	H	н
XA94	снз-	н	<u> </u>	HILL	н	н
XA95	СН3-	н	Н	HN,	Н	н
XA96	снз-	H	н		Н	Н
XA97	СН3-	н	н	QN >	н	Н
XA98	снз-	н	н	<b>1</b>	н	н
XA99	снз-	н	н	NO P	н .	н
XA100	снз-	н	н	SN .	н	н
XA101	снз-	Н	н	\$5,	н	н
XA102	снз-	н	н	1	н	н
XA103	снз-	н	н	O N	н	н
XA104	снз-	Н	н -		Н	Н
XA105	снз-	н	н	NS,	н	н
XA106	СН3-	н	н	S.	Н	н
XA107	СН3	H .	н	SY.	н	н
XA108	СН3-	н	н	NS,	Н	н
XA109	снэ-	н	н		нү	н
XA110	снз-	н	н		Н	អ .
XA111	снз-	н	н	N)	Н	н
XA112	СН3-	н	н		н	н
XA113	CH3-	н	н		н	н

No.	R1	R2	R3	R4	R5	R6
140.		<del>'`</del>	140	NEN	-	.,,,
XA114	CH3-	H	Н		Н	H
XA115	снз-	н .	Н		н	н
XA118	снз-	н	н	Ğ.	н	н
XA117	снз–	н	н.		н	Н
XA118	снз-	Н .	н		H·	Н
XA119	снз-	Н	н	,Ch	н	н
XA120	снз-	Н	H	Ţì	Н	Н
XA121	снз-	н	Н		н	H .
XA122	снз-	н	Н	O.	н	Н
XA123	СН3-	н	Н	Č:	н	Н
XA124	СН3	н	Н .	TO3	н	H
XA125	снз-	н	н	,CC	н	н
XA126	снз–	н	Н	Ţ3	н	н .
XA127 -	снз-	н	н		н	Н .
XA128	снз-	н	Н	OŞ	Н	н
XA129	снз-	н	н	Ċ;	Н	н
XA130	CH3-	н	н	TO3	н	н
XA131	снз-	н	н	,CI	н	н
XA132	снз-	н	H	Ţ?	н ,	н
- XA133	сна-	н	н	C'r	н	н
XA134	СН3-	н	н		н	н
XA135	снз-	н	н	TON	н	Н

No.	R1	R2	R3	R4	R5	R6
XA136	CH3-	н	н		H	Н
XA137	СН3-	н	н	Ţ,	н	H
XA138	CH3-	н	н		н	н
XA139	CH3-	Н	н	ŌŊ	Н	н
XA140	снз-	н	н	T N	Н	Н
XA141	СН3	н	н		н	Н
XA142	снз-	н	Н	Š <sub>o</sub>	Н	н
XA143	снз-	н .	н	'CL'	Н	Н
XA144	снз-	н.	Н	,CC	н	н
XA145	снз-	н	н	Ţ3	Н	H
XA146	снз-	н	н	O gr	н	н
XA147	СН3-	H 	н	Ū,	Н	н
XA148	снз–	Н	н	TON;	н	н
XA149 -	снз-	н	н	, CT	н	н
XA150	снз-	н	н	Ž <sup>s</sup>	н	н
XA151	снз-	H	н	C.	н .	н
XA152	снз-	Н	н	Ö	Н	Н
XA153	снз-	н	н	JOJ?	Н	н
XA154	CH3-	Н	н	,CC)	Н	н
XA155	СН3	н	н	Ğ.	Н	н
XA156	снз-	н	н	CT.	Н	н
XA157	CH3-	н	н	Ğ.	н	н

No.	R1	R2	R3	R4	R5	R6
	Ţ		1	<b>1</b>		1.00
XA158	снз-	Н	Н	1 Jan	Н	Н
XA159	снз-	Н	н ·	,CT\$ <sup>M</sup>	Н	н
XA160	снз-	Н	н	<u>Č</u> L3 <sub>4</sub>	Н	н
XA161	снз-	н	н	ر ا	Н	н
XA162	СН3-	н	Н		н	H
XA163	СН3-	Н	н	F O	н	н
XA164	снз-	Н	н	JO <sup>l</sup> ,	н	н
XA165	снз-	н	Н	G G	н	н
XA166	снз-	Н	Н	م کائے۔	н	н
XA167	снз-	Н	н	ر ا	Н	Н
XA168	снз-	н .	Н	Br O	Н	н
XA169	снз-	н	н	Br	н	н
XA170	снз-	н	H	Dir.	Н	н
XA171	снз-	н	Н	CHO.	н	н
XA172	снз-	н	н	H³C Q	н	н
XA173	снз-	н	H		н	н
XA174	СН3-	н ,	Н	CHIO O	н	Н
XA175	снз-	Н	Н	H3co	н	Н
XA176	снз-	Н	Н	H W J.	н	н
XA177	CH3-	н	Н	100	н	Н
XA178	СН3-	Н	Н	0 <sub>2</sub> N	н	н
XA179	СН3-	н	Н		Н	Н

No.	R1	R2	R3	R4	R5	Ine
140.	100	12	INS.	R4 GHO 人 人	lko –	R6
XA180	CH3-	К	н	O,	Н	н
XA181	снз-	н	н	HOSP	Н	н
XA182	снз-	Н	н		Н	н
XA183	снз-	н	н	NH O	н	Н
XA184	снз-	Н	н	HZN	н	Н
XA185	снз-	н	н	L.O.	Н	н
XA186	снз-	н	н	CN O	н	н
XA187	снз-	Н	н	NC P.	н	н
XA188	СН3-	н	н		H -	н
XA189	снз-	H	н		н	н
XA190	снз–	н	н	OO <sup>i</sup>	н	н
XA191	снз–	н	н	<u></u>	н	н
XA192	снз-	н	Н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA193 .	снз-	Н	Н	~\ <sup>2</sup> ,	н	н
XA194	снз–	н	Н	\rangle,	н	н
_ XA195	снз-	H	H.		н	н
XA196	снз–	н	Н	~\ <sup>2</sup> ,	н	Н
XA197	СН3-	н	Н	<b>,</b>	Н	н
XA198	снз–	Н	Н	~~ <sup>1</sup> ,	н	н
XA199	снз-	н	н	~~\ <sup>1</sup> ,	н	н
XA200	снз–	н	Н	~~~~°,	н	н
XA201	снз-	н	н .	√ <sup>2</sup> ,	н	н

No.	RI	R2.	R3	R4	R5	R6
XA202	CH3	н	н .		н	н
XA203	CH3-	H-	н	~ in	н	н
XA204	СН3-	н	н	74,	н	н
XA205	СН3	H³CQ_≻	н	н	н	н
XA206	СН3-	о н₃со <sup>л</sup> у	н	снз-	н .	н
XA207	СН3-	O H₃CO →	н	снзсн2-	H.	н
XA208	СН3	O H₃CO →	н	^ <u>`</u> `	Н	н
XA209	СН3-	H³CO_>.	н	Y	н	Н
XA210	СН3-	H³CO,≻	н .	<b>\\\</b>	Н	Н
,XA211	СН3-	H³CO, ≻	н	人、	Н	Н
XA212	снз-	H³CO,≻	н		н	н
XA213	снэ-	H³CO_>	н	丫	н	Н
XA214	снз-	H³CO,≻	н	~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н.	Н
XA215	снз–	H3CO 7	н	Y~~	н	Н
XA216	СН3-	H³CO, ⊁ Ö	Н	人」	н	н
XA217	снэ-	- H₃CO >	Н	Y	н	н
XA218	снз–	O H³CO√≻	н	<b>~~~</b> `	H	Н
XA219	снэ–	H³CO, λ. Ö	Н		Н	Н
XA220	снз-	H³CO_>\	н	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н
XA221	снз–	O H³CO,≻	н	Y~~~	н	Н
XA222	снз-	O H₃CO →	н	n-C8H17	н	Н
XA223	снз-	P³CO√≻	Н	L~~~	н	Н

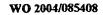
No.	R1	R2	R3	R4	R5	R6
XA224	снз-	H³CO, ≻	н	Q	н	Н
XA225	CH3-	H³CQ_≻ Ö	н		н	н
XA226	CH3-	H³CO, '\	н		Н	н
XA227	снз	H³CO, ≻	н	D;	н	Н
XA228	СН3	H³CO, ≻	н	$\Diamond$	н	н
XA229	снз-	H³CO_^\ Ö	н	D-1	н	н
XA230	снз-	O H₃CO >	н	$\bigcirc$ $\dashv$	н .	н
XA231	снз-	O H₃CO y	н	$\bigcirc$ $\dashv$	Н	н
XA232	снз-	H³CO, ≻	Н		н	н
XA233	CH3-	D H₃CO >	Н		н .	н
XA234	снз-	H³CO, ≻	н	<b>_</b>	н	Н
XA235	снз-	O H₃CO →	н	F-()-1	н	н
XA236	CH3-	H₃CO →	н	CI →	н	Н
XA237	снз-	H³CO, ≻	н	CI	н	н
XA238	CH3-	H³CO_≻	н	c⊢ <b>(</b> )~{	н	н
XA239	CH3-	H³CO, ≻	. н	Br →{	н.	н
XA240	снз-	H₃CO >	н	Br.	н .	н
XA241	снз-	H³CO_≻	н	Br-{}-{	Н	н
XA242	СН3-	H₃CO >	н	CH₃	н	н
XA243	снз-	H₃CO >	н	H₃C →	н	н
XA244	СН3-	H <sub>3</sub> CO >	н	H <sub>3</sub> C-{	н	н
XA245	снз-	H <sub>3</sub> CO >	н	C <sub>2</sub> H <sub>5</sub> -{}-{	н	н

•		_
		(
		١.

No.	R1	R2	R3	R4	R5	R6
XA246	снз-	H³CO,≻	Н	n-C <sub>3</sub> H <sub>7</sub> -	Н	H
XA247	снз-	H³CO_≻	н	n-C <sub>4</sub> H <sub>9</sub> -	н	н
XA248	снз-	H³CO,≻	н	OCH <sub>3</sub>	н	Н
XA249	снз-	O H³CO,≻	н	H <sub>3</sub> CO	н	н
XA250	снз-	H₃CO →	н	H₃CO-{_}-{	н	н
XA251	снз-	H³CO, ≻	н	C <sub>2</sub> H <sub>5</sub> O-{}-{	н	н
XA252	снз-	O H₃CO <sup>*</sup> }⁄	н	n-C <sub>3</sub> H <sub>7</sub> O-⟨}-	н	Н
XA253	СН3-	O H₃CO →	н	n-C₄H <sub>9</sub> O-⟨}{	н	н
XA254	снз-	H³CO, ≻ Ö	Н	NO <sub>2</sub>	н	н
XA255	СН3	O H₃CO >	н	O <sub>2</sub> N	н	н
XA256	снэ-	H³CO, ≻	Н	02N-	н	н
XA257	снз-	O H₃CO ≻	н	CN →	н	н
XA258	CH3-	H³CO, ≻	н	NC	н	н
XA259	СН3-	O H₃CO ≻	н	NC-()-{	Н	Н
XA260	СН3-	O H₃CO →	н	NMe <sub>2</sub>	н	н
XA261	снз-	O H₃CO >	н	Me <sub>2</sub> N	н	н
XA262	снэ–	H³CO, ≻	н	Me <sub>2</sub> N-{}	н	н
XA263	снз-	H₃CO <sup>^</sup> >	н	OP.	н	Н
XA264	СН3-	H³CO_>	н	OO'	н	н
XA265	снз-	H³CO_>	н	O's	н	н
XA266	СН3~	O H₃CO →	н	Q.	н	Н
XA267	снз-	O H³CO ≻	н	OJ,	н	н

# PCT/JP2004/004320

No.	R1	R2	R3	R4	R5	R6
XA268	снэ-	H³CO_≻	Н	R4 O ,,,	н	н
XA269	снз-	O H³CO,≻	н	Ŷ,	н	н
XA270	снз–	O C₂H₅O ≻	н	н	н	н
XA271	СН3	C₂H₅O >	н	снз-	н	н
XA272	снз-	C₂H₅O →	Н	снзсн2-	Н	н
XA273	снз-	O C₂H₅O ≻	H	^	н	н
XA274	снз-	C <sub>2</sub> H <sub>5</sub> O <sup>™</sup> ≻	Н	Υ	н	н
XA275	снз-	O C₂H₅O ≻	Н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA276	снз–	C₂H₅O <sup>™</sup> ≻	Н	从	н	н
XA277	снз-	C₂H₅O ≻	H		н	н
XA278	СН3-	O C₂H₅O →	н	7	Н	н
XA279	снз–	C <sub>2</sub> H <sub>5</sub> O >	н		н	н
XA280	снз–	C <sub>2</sub> H₅O ≻	Ĥ	Y~	н	н
XA281	снз–	C2H2O√>	Н .	人 ス	н	н
XA282	снз-	C <sup>2</sup> H <sup>2</sup> O, 2,	н	7	н	н
XA283	снэ-	· C <sup>2</sup> H <sup>2</sup> O , \	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA284	снз-	C₂H₅O <sup>™</sup> >	н	<u></u>	н	н
XA285	СН3	O C₂H₅O ̈́≻	Н		н	Н
XA286	СН3-	C⁵H²O, ≻	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
XA287	снз-	O C₂H₅O >≻	Н	n-C8H17-	н	н
XA288	снз-	C <sub>2</sub> H <sub>5</sub> O <sup>1</sup> >r	н	L	Н	H
XA289	снз-	C2H2O >	Н	Qu	н	н



No.	R1	R2	R3	R4	R5	R6
XA290	снз-	O C <sub>2</sub> H <sub>5</sub> O √ ≻	н	0	н	Н
XA291	CH3-	C <sup>2</sup> H <sup>2</sup> O <sub>4</sub> > O	н	Qir	н	Н
XA292	снз-	O C <sub>2</sub> H <sub>5</sub> O /	н	<b>D</b>	н	н
XA293	снз-	O C <sub>2</sub> H₅O √	н	$\Diamond$ -1	Н	н
XA294	CH3-	C <sub>2</sub> H <sub>5</sub> O <sup>+</sup> >	Н	$\bigcirc$	н	Н
XA295	CH3-	C <sub>2</sub> H <sub>5</sub> O >	н	$\bigcirc$ $\dashv$	н	Н
XA296	СН3	O C₂H₅O ,≻	н	$\bigcirc$	н	н
XA297	снз-	C₂H₅O <sup>Ĩ</sup> →	н		Н	Н
XA298	снз-	C²H²OД³	н	<b>\( \)</b>	Н	н
XA299	снз-	C <sup>5</sup> H <sup>2</sup> O , >,	Н		н	н
XA300	снз-	C₂H₅O →	н	F-()-1	н	н
XA301	снз-	C <sub>2</sub> H <sub>5</sub> O >	н	CI CI	н	н
XA302	снз-	C <sup>2</sup> H²O√≻	н	CI	н	н
XA303	снз-	C <sub>2</sub> H <sub>5</sub> O y	Н	CH	н	н
XA304	CH3-	C <sub>2</sub> H <sub>5</sub> O <sup>1</sup>	н	Br	н	н
XA305	снз-	C₂H₅O >	H	Br.	н	н
XA306	CH3-	C₂H₅O y	Н	Br-{}-{	Н	н
XA307	СН3-	C <sub>2</sub> H <sub>5</sub> O <sup>1</sup> >	н	CH <sub>3</sub>	н	н
XA308	снз-	C <sub>2</sub> H₅O >	н .	H <sub>3</sub> C	н	н
XA309	снз-	C₂H₅O >	Н	H <sub>3</sub> C- <b>(</b> )	Н	н
XA310	СН3-	C2H2O_2	н		н	н
XA311	снз-	C <sub>2</sub> H <sub>5</sub> O y	н	ᠬᢗ₃Hᠨ᠆⟨█ <mark>〉</mark> ᢇ┤	н	н

No.	R1	R2	R3	R4	R5	R6
XA312	снз-	C <sub>2</sub> H <sub>5</sub> O >	н	n-C <sub>4</sub> H <sub>9</sub> -	н	н
XA313	СН3-	C <sub>2</sub> H <sub>5</sub> O <sup>2</sup> /	н	OCH₃	н	н
XA314	снз-	C <sub>2</sub> H <sub>5</sub> O >	н	H <sub>3</sub> CO	н	Н
XA315	снз-	C <sub>2</sub> H <sub>5</sub> O >	н	H <sub>3</sub> CO-{}-{	Н	н
XA316	CH3-	C <sub>2</sub> H <sub>5</sub> O <sup>2</sup> /	н		н	н
XA317	СН3	O. C₂H₅O >⁄	н	n-C <sub>3</sub> H <sub>7</sub> O-	н	н
XA318	снз-	C <sub>2</sub> H <sub>5</sub> O /	н	n-C <sub>4</sub> H <sub>9</sub> O-	н	н
XA319	CH3-	C <sub>2</sub> H <sub>5</sub> O 7	н	NO <sub>2</sub>	н	н
XA320	снз-	O C₂H₅O →	н	O <sub>2</sub> N	н	Н
XA321	СН3-	C <sub>2</sub> H <sub>5</sub> O >	н	02N-()-(	н	н
XA322	снз–	C <sub>2</sub> H <sub>5</sub> O <sup>2</sup> /	н	CN	н	н
XA323	снз-	C <sub>2</sub> H <sub>5</sub> O y	Н	NC	н	н
XA324	снэ–	O C₂H₅O ≻	н	NC-()-1	н -	н
XA325	снз-	C <sub>2</sub> H <sub>5</sub> O >	н	NMe <sub>2</sub>	н	н
XA326	снз-	C⁵H²O, ∖.	н	Me <sub>2</sub> N	н	Н
XA327	снз-	C <sub>2</sub> H <sub>5</sub> O /	. н	Me <sub>2</sub> N-{}-{	н.	н
XA328	снз-	C <sub>2</sub> H <sub>5</sub> O √y	н		н	н
XA329	CH3-	C <sub>2</sub> H <sub>5</sub> O >	н	OO'	н	н
XA330	СН3-	C2H2O_>	н	رث ا	н	н
XA331	снз-	C₂H₅O →	Н	Qi,	н	н
XA332	снз-	C <sub>2</sub> H <sub>5</sub> O <sup>1</sup> >	н	OO',	н	н
XA333	СН3	C <sub>2</sub> H <sub>5</sub> O >	н	9,	н	н

No.	R1	R2	R3	R4	R5	R6
				<u>,</u> ,		
XA334	СН3-	°C2H²O√≻	H	<b>~</b>	н	H
XA335	снз–	СН3-	Н	н	н	Н
XA336	снз-	снзсн2-	H	н	н	н
XA337	СН3-	<u> </u>	Н	н	н	H
XA338	снз-	$\rightarrow$	н	н	н	н Н
XA339	СН3	<b>\</b> \\\	Н	Н	н	н
XA340	снз-	人人	Н	н	Н	н
XA341	снз–		Н	н .	н	Н
XA342	снз–	丫	Н	н	н	н
XA343	СН3	^^\	Н	н	н	н
XA344	СН3	<b>Y</b>	н	H .	н	н
XA345	СН3	人、	н	н	Н	н
XA346	снз	$\gamma$	н	Н	н .	H
XA347	СН3-	~~^\	н	н	н .	н
XA348	снз-	L~~	н	H	н	н
XA349	СН3-	^^^\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	H	н	н	н
XA350	снз-	Y~~~	 Н	Н	н .	Н
XA351	снз-	n-C8H17-	н	н _	н	Н
XA352	снз–	L~~~	н	Н	н	н
XA353	СН3-	Qu.	н	н .	н	н
XA354	снз-	0~~	н	н	н	н
XA355	снз-	Q	н	н	Н	н

No.	R1	R2	R3	R4	R5	R6
XA356	снз-	<b>▷</b> →	н	н	Н	н
XA357	снз-	$\Diamond$	н	н	н	н
XA358	снз-	$\bigcirc$	Н	н	н	н
XA359	снз–	$\bigcirc$ $\dashv$	н	н	н	н
XA360	снз-	$\bigcirc$	н	н	Н	Н
XA361	СН3-		Н	н	H ·	н
XA362	СН3-		Н	н .	Н	н
XA363	СН3-	<u>⟨``</u> }{	Н	н	н	н
XA364	CH3-		Н	Н	н	н
XA365	СН3-		Н	н	Н	н
XA366	СН3-	F-C)-i	Н	Н	Н	н
XA367	снз–	F-(-)(	Н	н	н	н
XA368	снз-	F{>ı{	н	н .	Н	н
XA369	снз-	CI	н	н	н	н
XA370	снз–	CI	н	н	н	H
XA371	CH3-	c <del>-(_)-</del>	н	н	н	н
XA372	СН3-	c⊢( <b>&gt;</b> -{	н	н	Н	н
XA373	снз–	CI()···-(	Н	H .	н	н
XA374	CH3	Br C≻⊰	Н	н	н	н
XA375	СН3-	Br.	н	Н	н	Н
XA376	СН3	Br-{}-{	н	н	Н	н
XA377	СН3	Br—{}	н	н	Н	н

<del></del>	154	ina .			loe .	50
No.		R2	R3	R4	R5	R6
XA378	CH3-	Br-《》····{	н	н	н	H .
XA379	снэ-	<b>_</b>	н	н	н	н
XA380	снз-		н	н	н	н
XA381	снз-		Н	н	н	H
XA382	снз-	CH <sub>3</sub>	Н	Н	н	Н
XA383	снз-	H <sub>3</sub> C	н	н	Н	Н.
XA384	снз-	H <sub>3</sub> C-{_}-{	Н	н	н	Н
XA385	снз-	C₂H₅- <b>⟨</b> }{	н	н	н	Н
XA386	снз-	n-C <sub>3</sub> H <sub>7</sub> -	н	н	н	н
XA387	снз–	n-C₄H <sub>9</sub> -∕	н	Н	н	н
XA388	снз-	OH OH	н	н .	Н	н
XA389	снз-	HO	н	Н	Н	н ,
XA390	снз-	но-{}-} .	Н	н .	н	н
XA391	снз-	OCH <sub>3</sub> .	н	н	н	н
XA392	СН3-	H₃CO ☐	Н	н	н	н
XA393	снз–	H³CO-{_}_	H	н	н .	н
XA394	снз-	H3CO-()-(	н	Н	Н	н
XA395	снз-	H <sub>3</sub> CO-{_}\\	н	Н	Н	н
XA396	снз-	OC <sub>2</sub> H <sub>5</sub>	Н	Н	н	н
XA397	снз–	C2H2O	н	Н	Н	н
XA398	СН3-	C <sub>2</sub> H <sub>5</sub> O-{_}-	н	н	н	н
XA399	СН3	n-C <sub>3</sub> H <sub>7</sub> O-	Н	н	н	н

No.	RI	R2	R3	R4	R5	R6
XA400	CH3-	n-C <sub>4</sub> H <sub>9</sub> O-{}{	н	н	н	н
XA401	снз-	NO <sub>2</sub>	н	н	н	н
XA402	CH3-	O <sub>2</sub> N	н	н	н	н
XA403	снз-	O <sub>2</sub> N-{}	н	н	н	н
XA404	CH3-	CN ∰-¦	н	н	н	н
XA405	CH3-	NC.	Н	н	Н	н
XA406	СН3-	NC-	Н	н	Н	Н
XA407	снз-	CF₃	н	Н	н	н
XA408	снз-	F₃C	Н	н	н	н
XA409	CH3-	F <sub>3</sub> C-{	н	н	н	н
XA410	снз-		Н	н	Н	н
XA411	снз-	HOOC	Н	н	н	н
XA412	снз–	H00C-{\rightarrow}-{	н	н	н	н
XA413	снз-	CO <sub>2</sub> Me	Н	н .	н	н
XA414	CH3-	MeO <sub>2</sub> C	н	н	н	н
XA415	снз-	MeO <sub>2</sub> C-{	н	н	н	н
XA416	СН3-		н	н	н _	н
XA417	СН3-	EtO <sub>2</sub> C	н	н 	н	н
XA418	снз–	EtO <sub>2</sub> C-{}-{	н	Н	н	H
XA419	снз–	SMe	н	н	н	н
XA420	снз-	MeS	Н	н	н	Н
XA421	снз-	MeS-{	н	Н	Н	н .

No.	R1	R2	R3	R4	R5	R6
XA422	снз-	SO₂Me	н	н	н	Н
XA423	СН3-	MeO₂S	н	н	H	н
XA424	снз-	MeO₂S-{_}_	н	н	н	н
XA425	снз-	NH <sub>2</sub>	н	н	н	н
XA426	СН3	H <sub>2</sub> N	н	Н	н	н
XA427	CH3	H <sub>2</sub> N-{}	н	н	Н	н
XA428	снз-	NMe <sub>2</sub>	н	н	н	н
XA429	СН3	Me₂N	н	н	н	Н
XA430	СН3-	Me <sub>2</sub> N-{}	Н	Н	Н	Н
XA431	снз-		н,	н	н	н
XA432	СН3-	CHQ)	н	н	н	н
XA433	снз-	CH-C)-1	Н	н	н	H -
XA434	СН3		Н	н	Н	н
XA435	CH3-		н	Н	Н	н
XA436	снз-	O+⊘-₁	н	Н	н	н
XA437	снз-		н	н	н	Н
XA438	CH3-	<b>€</b>	Н	н	н	н
XA439	снз-		н	н	Н	н
XA440	СН3-	H³CN N-{}	Н	Н	н	н
XA441	снз-	H3CN (N-{\( \) \	Н	н	н	н
XA442	снз-	H3CN_N-{}-{	н	Н	н	н
XA443	снз-	H₃C CH₃	н	Н	н	Н

No.	R1	R2	R3	R4	R5	R6
XA444	CH3-	CH <sub>3</sub> H <sub>3</sub> C-{}}	Н	н	Н	Н
XA445	СН3-	Сн₃ (∑)-; н₃с	н	н	н.	Н
XA446	CH3-	CH₃ CH₃	н	н	н	н
XA447	СН3-	H <sub>3</sub> C-(_)-{	Н	Н	Н	Н
XA448	снз-	H₃C H₃C	Н	H	Н	н
XA449	снз-	F_F	Н	н .	Н	Н
XA450	снз-	F-G	Н	н	н	Н
XA451	CH3-		Н	н	н	Н
XA452	СН3	<b>⊕</b>	H	н .	н	Н
XA453	снз-	F-{\rightarrow}	н	н	н	Н
XA454	снз-	F	н	Н	н	H
XA455	СН3-		н	н	н	Н
XA456	снз–	a-{};	н	н	Н	н
XA457	СН3-	l ~'	Н	н	н	Н
XA458	снз-		Н	н	н	Н
XA459	снз-	a— d—	н	н	н	н

No.	Ri	R2	R3	R4	R5	R6
XA460	СН3-	a - a	н	н	н	H
XA461	СН3-	H₃CQ_OCH₃	Н	н	н	н
XA462	СН3-	H³∞-{_}-\ OCH3	н	Н	Н	Н
XA463	CH3~	OCH <sub>3</sub> ———————————————————————————————————	Н	н	H.	Н
XA464	снз–	OCH <sub>3</sub>	н	Н	н	Н
XA465	СН3-	H3CO	н	Н	Н	н

No.	R1	R2	R3	R4	R5	Inc
- 1,52	<del>                                     </del>		1100	<del>   </del>	(R5	R6
XA466	снз-	H <sub>2</sub> CO	н	Н	Н	Н
XA467	снз–	F_OCH₃	н	н	Н	н
XA468	снз-	OCH <sub>3</sub>	н	Н	Н	Н
XA469	снз-	OCH <sub>3</sub>	н	Н	Н	н .
XA470	СН3-	OCH <sub>3</sub> F—⟨⟩⊪{	Н	H	Н	н
XA471	снз-	OCH <sub>3</sub> F	Н	Н	н	н
XA472	CH3-	OCH <sub>3</sub>	Н	Н	н	Н
XA473	CH3-	H₃CQ F—	н	н	н	н
XA474	снз-	H₃CQ F	Н	н.	н	Н
XA475	снз–	H³CO_È	Н	Н	н	Н
XA476	СН3	H <sub>3</sub> CO-{\bigs_}}	н	Н	Н	н
XA477	снз-	H <sub>2</sub> CO F	н	Н	Н	н
XA478	СНЗ-		н	Н	Н	Н
XA479	СН3	C OCH3	Н	Н	н	H
XA480	снз-		н	Н	Н	Н
XA481	CH3-	OCH₃ CI	н	Н	Н	Н

No.	R1	R2	R3	R4	R5	R6
XA482	СН3-	C)→ OCH3	н	н	Н	Н
XA483	снз-	H <sub>3</sub> CO	н	н	н	Н
XA484	СН3-	H³CO →	н	н	Н	н
XA485	снз-	H³CO CI	н	н	н	н
XA486	снз-	H₃CO-{	Н	н	н	н
XA487	СН3-	CI H₃CO	н	н	н	Н

No.	RI	R2	R3	R4	R5	R6
XA488	СН3-	CI H <sub>3</sub> CO-{}-{	Н	Н	н	н
XA489	снз-	F_CH <sub>3</sub>	н	н	н	н
XA490	СН3-	CH <sub>3</sub> F-{∑}-;	Н	н	н	Н
XA491	СН3-	CH <sub>3</sub> F	Н	Н	н	Н
XA492	снз-	CH₃ F	н	н	н	н
XA493	CH3-	H <sub>3</sub> C F—{}-{	Н	н	н	Н
XA494	СН3-	H <sub>3</sub> C	н	н	н	н
XA495	СН3-	H₃C _ F	Н	н	н	н
XA496	СН3	H₃C-⟨¯¯¯⟩}	н .	н	н	Н
XA497	снз-	H <sub>3</sub> C	н	Н	Н	н
XA498	снз-	H³C-⟨}-\	Н	Н	Н	н
XA499	СН3-	\ \\ \'	Н	н	Н	н
XA500	CH3-	OCH₃ Br—⟨□}→	H	н	Н	Н
XA501	СН3-	OCH <sub>3</sub> ⇒	Н	н	н	н
XA502	СН3-	OCH <sub>3</sub> Br	H	н	Н	Н
XA503	СН3	H <sub>3</sub> CQ Br→{}	н	н	н	Н

No.	R1	R2	R3	R4	R5	R6
XA504	снз-	H <sub>3</sub> CQ Br	н	Н	н	н
XA505	CH3-	H <sub>3</sub> CO_Br	н	Н	н	Н
XA506	СН3-	H₃CO-⟨¯¯⟩¦	н	н	Н	Н
XA507	снз	Br H₃CO	н	н	н	н
XA508	СН3-	H <sub>3</sub> CO	н	н	н	Н
XA509	СН3-	()4-(_) H³co >	н	н .	н	Н

No.	R1	R2	R3	R4	R5	R6
	<u> </u>	OCH <sub>3</sub>	140	174	iko –	INV
XA510	снз–		н	н	н	Н
XA511	CH3-	Cn-C}-och³	н	Н	Н	Н
XA512	СН3-	H³CO →	Н	Н	н	н
XA513	СН3-	H₃CO ○N-(□);	н	н	н	н
ХА514	снз-	Ch Sylvanian Character Ch	Н	н	н	н
XA515	СН3-	F—(S)+ F	н	н	Н	н
XA516	СН3-	OCH₃ F————————————————————————————————————	Н	Н.	Н	н
XA517	снз-	H₃CO-{\}_{F}	Н	н	н	Н
XA518	СН3-	OCH <sub>3</sub> F-C	Н	н	H	Н
XA519	СН3-	OCH <sub>3</sub> OCH <sub>3</sub>	н	Н	н	Н
XA520	СН3	ci—(∑); a	Н	н	Н	н
XA521	снз-	OCH <sub>3</sub> CI—(C)—( CI	Н	н	Н	Н
XA522	CH3-	H₃co-<_}; a	Н	н	н	н
XA523	Снз-		Н	н	н	н

No.	R1	R2	R3	R4	IDE	155
XA524	СН3-	OCH <sup>3</sup>	Н	Н	R5 H	R6 H
XA525	снз-	OCH <sub>3</sub>	Н	н	н	н
XA526	СН3-	H <sub>3</sub> CO	н	Н	Н	н
XA527	СН3-	H3CO-{\}-{\}-{	Н	н .	н	Н
XA528	снз-	OCH <sub>3</sub> ;\	н	н	н	н
XA529	СН3-	H-00	Н	н	н	Н
XA530	СН3-	H3CO-{\rightarrow}^\tag{\chi}	Н	н	н	Н
XA531	СН3-	OCH <sub>3</sub>	Н	H	Н	н

No.	R1	R2	R3	R4	R5	R6
XA532	СН3	H-CQ	Н	н	Н	Н
XA533	снз-	H <sub>3</sub> CO-{	н	н	н	н
XA534	СН3-	<b>₫</b>	Н	н	н	н
XA535	СН3-		н	н	н	Н
XA536	СН3-	F-(>-(>-1)	н	н	Н	н
XA537	СН3-		Н	н	Н	н
XA538	снз-		Н	Н	н	Н
XA539	снз-		Н	Н	н	н
XA540	снз–	ď.	н	Н	н	н
XA541	снз-	\$\_\( \infty\)	Н .	Н	Н	н
XA542	CH3-		н	н .	н	н
XA543	снэ-		н	Н -	Н	H.
XA544	снз–	CC,	Н	Н	н	н
XA545	снз	E T	Н	н	Н	н
XA546	СН3-	HN	Н	Н	н	н
XA547	CH3-		Н	Н	н	н

No.	R1	R2	R3	R4	R5	R6
XA548	СН3-	5	н	Н	н	н
XA549	СН3-	(S)	Н	Н	H	н
XA550	СН3-	\$7,	Н	н	н	Н
XA551	СН3-	HNN	Н	н	Н	н
XA552	СН3-	HN	Н	н	н	Н
XA553	СН3	HN.	н	н	н	н

No.	R1	R2	R3	R4	R5	R6
		FN ·				
XA554	снз-	LZ.	Н	Н	Н	Н
XA555	снз–	d√~	н	н	H	н
XA556	снз–	05	н	н	н	н
XA557	снз-	MO	н	н	н	н
XA558	снз-	S <sub>N</sub>	н	н	н	Н
XA559	СН3-	\$5,	н	н	н	Н
XA560	снз–	N-S V->	Н	н	н	н
XA561	СН3-	€N,	Н	н	Н	н
XA562	снз–	\$\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	н .	н	н
XA563	СН3-	N.S.,	Н	н	Н	н
XA564	снз-	S,	н	н	H	н
XA565	СН3-	SY,	н	Н	Н	н
XA566	снз-	N,	Н	Н	н	н
XA567	СН3-	€N-1	н.	Н	Н	н
XA568	снз-	<b>₩</b>	н	н	н	н
XA569	снз-	<b>!</b> ○→1	н	н .	H	Н
XA570	снз-		н	н	н	н
XA571	СН3-		Н	н	Н	н
XA572	СН3-	N-	Н	н	н	н
XA573	снз-		н	н	н	н
XA574	снз-		Н	н	н	н
XA575	снз-		н	н	н	н

No.	R1	R2 ·	R3	R4	R5	R6
XA576	снз-	(C)	н	н	н	н
XA577	снз	,CT	н	н	н	н
XA578	снз-	Çî,	н	н	н	Н
XA579	СН3-	CD;	н	н	н	н
XA580	снз-	O.	н	н	н	н
XA581	снз-	Ö	Н	н	н	н
XA582	снз-	TOP	н	н	н	н
XA583	снз-	,CC)	н	Н	Н	н
XA584	снз-	Q:	н	н	н	Н
XA585	снз-	CT\$+	н	н	Н	н
XA586	снз-		. Н	н	Н	н
XA587	снз-	(C)	н	Н	н	Н
XA588	снз–	TI?	н	н	Н	н
XA589	снз-	,CIŞ	н	Н	н	н
XA590	снз-	ÇŢ\$	н	н	. н	н
XA591	CH3-	CÎ	н	н	н	н
XA592	снз-	T H	н	н	н	Н
XA593	снз-	TON THE PROPERTY OF THE PROPER	н	Н	н	н
XA594	CH3-	,CTh	н	н	н	Н
XA595	снз-	Ţ'n.	н	Н	н	Н
XA596	CH3-	CIN-1	Н	Н	Н	н
XA597	снз-	Ţ,»	н	н	н	н

	lot -	los	200	In a	lDE .	ne l
No.	R1	R2	R3	R4	R5	R6
XA598	снз–		н	н	Н	H
XA599	снз-		Н	Н	Н	H
XA600	снз–	Ţ <sub>N</sub>	н	н	н	H
XA601	снз-		Н	н	н	н
XA602	СН3	4 (I)	н	Н	н	H
XA603	CH3-		н	н	н	Н
XA604	снз-	Ž <sup>N</sup>	Н	Н	н	Н
XA605	СН3-	J.N.	н	н	Н	Н
XA606 .	СН3	'Cly	н	н	н	н
XA607	СН3	√O's	н	н	н	Н
XA608	снз-	Q <sup>N</sup> <sub>s</sub>	н	Н .	н	H
XA609	снз-		н	н	н	Н
XA610	снз–		н	Н	Н	Н
XA611	снз-	TO TO	н	н	H	Н
XA612	снз-	,CT%	н	Н	Н	н
XA613	снз-	Ţ,	н	н	Н	н
XA614	снз-		н	н	Н	н
XA615	снз–	Ţ,	н	H	Н	н
XA616	снз-	TOP .	н	н	н	Н
XA617	CH3-	,CT3h	н	н	Н	н
XA618	снз-	Ţ,	н	Н	Н	н
XA619	снз-	Ţ,	н	н	н ,	н



No.	Rt	R2	R3	R4	R5	R6
NO.			<u></u>	1/4	iw.	110
XA620	снз-	,CC),	н	Н	H	Н
XA621	СН3-	<b>TOS</b>	Н	н	Н	н
XA622	СН3-	O	н	Н	н	н
XA623	CH3-	снз-	Н	снз	н	н
XA624	СН3-	СН3СН2-	Н	снз	н	н
XA625	CH3-	<b>^</b> \`	н	снз	Н	н
XA626	СН3-	丫	Н	снз	н	н
XA627	снз⊸	<b>✓</b> ✓ .	Н	СНЗ	н	Н
XA628	снз-	人人.	Н	снз	Н	Н
XA629	снз-	$\uparrow \uparrow$	Н	снз	Н	Н
XA630	снз-	丫	Н	снз	н	Н
XA631	СН3-	~~`\	н	снз	Н	Н
XA632	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	снз	Н	н
XA633	СН3-	<u> </u>	н	снз	н	н
XA634	снз-	7	н	снз	н	н
XA635	снз-	<b>~~~</b>	н	снз	н	н
XA636	СН3-	L~~	н	СНЗ	н	н
XA637	снз-	~~~``	н	снз	н	н
· XA638	снз-	~~~``	н	СНЗ	н .	Н
XA639	снз-	n-C8H17-	н	СНЗ	н	н
XA640	СН3-	L~~~	н	снз	н	н
XA641	CH3-		н	СНЗ	н	Н

No.	R1	R2	R3	R4	R5	R6
XA642	снз-	$\sim \sim$	Н	СНЗ		н
XA643	снз-		H	СНЗ	Н	Н
XA644	снз-	$\triangleright$	Н	CH3	Н	н
XA645	снз-	$\Diamond$	н	СНЗ	Н	н
XA646	снз-	$\bigcirc$	H	CH3	н	н Н
XA647	снз-		н	СНЗ	н	н
XA648	снз-	$\bigcirc$	н	СНЗ	н	Н
XA649	снз–		Н	СНЗ	н	н
XA650	снз-	$\bigcirc$	н	СН3	н	н
XA651	снз-	<b>⊘</b> m4	н	СН3	н .	н
XA652	СН3-	F .	Н	снз	н	Н
XA653	СН3		H	сн3	н	Н
XA654	снз-	F-()-1	Н	снз	н	н
XA655 -	снз–	F-(>-1	н	снз	Н	н
XA656	снз-	F{>ı::-{	н	снз	Н	н
XA657	снз-	CI	н	сн3	Н	н
XA658	СН3-	CI	н	СНЗ	н	н
XA659	СН3-	cl-(	н	снз	н	н
XA660	СН3	c⊢ <b>(</b> )~-	н	снз	н .	Н
XA661	снз-	CI—(	н	снз	н	Н
XA662	СН3-	Br ←	н	СНЗ	н	Н
XA663	CH3-	Br.	н	снз	н	н





·	<u> </u>					
No.	R1	R2	R3	R4	R5	R6
XA664 .	снз-	Br-∕}-{	н	снз	н	н
XA665	CH3-	Br-{}	н	СНЗ	н	Н
XA666	снз-	Br—{	н	СНЗ	н	Н
XA667	снз-		H.	СНЗ	н	н
XA668	снз-		н	CH3	н •	н
XA669	СН3-		н	СНЗ	н	н
XA670	снз-	CH₃	н	СНЗ	Н	н
XA671	снз-	H <sub>3</sub> C	н	снз	н	н
XA672	снз-	H <sub>3</sub> C-	н	снз	н	н
XA673	снз-	. C <sub>2</sub> H <sub>5</sub> —{	Н	СНЗ	н	н
XA674	снз-	n-C₃H <sub>7</sub> {_}}{	н	СНЗ	н	н
XA675	снз-	n-C <sub>4</sub> H <sub>9</sub> {}-{	н	СНЗ	н	Н
XA676	снз-	ОН	Н	снз	н	н
XA677	СН3-	HO HO	H	СНЗ	н	н
XA678	снз-	HO-{}-}	н	снз	н	н
XA679	снз-	OCH₃	н	СНЗ	н	н
XA680	снз-	H <sub>3</sub> CO	н	CH3	н	н .
XA681	СН3-	H³CO- <b>⟨</b> }−4	н .	СНЗ	н	н
XA682	снз-	H3CO-{}-	н	СНЗ	н	н
XA683	СН3	H <sub>3</sub> CO-{\rightarrow}m{\rightarrow}m{\rightarrow}	н	СНЗ	н	н .
XA684	снз-	OC <sub>2</sub> H <sub>5</sub>	н	СНЗ	н	н
XA685	СНЗ-	C <sub>2</sub> H <sub>5</sub> O	н	снз	н	н

No.	R1	R2	R3	R4	R5	R6
No.	IKI		K3	K4	rcs	100
XA686	СН3-	C <sub>2</sub> H <sub>5</sub> O-(){	Н	СНЗ	Н	н
XA687	CH3-	n-C₃H <sub>7</sub> O-⟨}-{	Н	снз	Н	н
XA688	CH3	n-C <sub>4</sub> H <sub>9</sub> O-	н	снз	н	н
XA689	снз-	NO <sub>2</sub>	Н	снз	Н	н
XA690	СН3	O <sub>2</sub> N	н	снз	Н	Н
XA691	снз-	O <sub>2</sub> N-{}	н	СНЗ	н	н
XA692	снз-	CN	Н	снз	н	н
XA693	CH3-	NC ————————————————————————————————————	н	снз	Н	Н
XA694	СН3-	NC-(	н	снз	н	н
XA695	СН3-	CF₃	Н	СНЗ	н	Н
XA696	СН3-	F <sub>3</sub> C	Н	СНЗ	н	н
XA697	СН3	F <sub>3</sub> C-{}-{	н	СНЗ	н	н
XA698	СН3-	СООН	Н	СНЗ	н	н
XA699	СН3-	HOOC	н	снз	н	н
XA700	CH3-	H00C-{\rightarrow}-4	н	снз	н	н
XA701	снз-	CO₂Me	н	снз	н	н
XA702	снз	MeO <sub>2</sub> C	н	снз	н	н
XA703	снз-	MeO <sub>2</sub> C-	н	снз	н.	н
XA704	снз-	CO₂Et	Н	снз	Н	н
XA705	снз-	EtO <sub>2</sub> C	н	СНЗ	Н	Н
XA706	снз-	EtO <sub>2</sub> C-	н	СНЗ	Н	н
XA707	снз-	SMe	н	снз	н	н

No.	RI	R2	R3	R4	R5	R6
XA708	снз–	MeS	н	снз	Н	Н
XA709	СН3-	MeS-{}-{	н	снз	н	Н
XA710	снз-	SO₂Me	н	снз	Н	н
XA711	СН3-	MeO <sub>2</sub> S	н	снз	Н	Н
XA712	снз	MeO <sub>2</sub> S-	н	снз	Н	Н
XA713	снз-	NH₂	н	снз	Н	н
XA714	СН3-	H <sub>2</sub> N	н	СНЗ	Н	н
XA715	снз-	H <sub>2</sub> N-{}-{	н	снз	н	н
XA716	снз-	NMe <sub>2</sub>	н	снз	н	н
XA717	СН3-	Me <sub>2</sub> N	н	снз	н	н
XA718	снз-	Me <sub>2</sub> N-{	н	снз	H ·	н
XA719	снз-	Or D	н	снз	н	н
XA720	СН3-	OHQ.	н	снз	Н	н
XA721	CH3		н	снз	Н	н
XA722	СН3-		н	снз	Н	н
XA723	снз-		н	СНЗ	Н	н
XA724	СН3		н	снз	Н	н
XA725	снз-		н	снз	н	Н
XA726	снз-		н	СНЗ	н	н
XA727	снз-		н	снз	н	н
XA728	СН3-	H3CN N-	н	снз	н	н
XA729	снз-	H3CN N-	н	снз	н	н

No.	RI	R2	R3	R4	R5	R6
XA730	CH3-	H³CN_V-{_}-{	н	снз	н	н
XA731	снз-	H₃C_CH₃	Н	снз	н	н
XA732	снз-	CH <sub>3</sub>	н	СНЗ	н	н
XA733	снз-	CH <sub>3</sub> CH <sub>3</sub> H <sub>3</sub> C	H	СНЗ	н	Н
XA734	снз-	CH₃ CH₃	Н	СНЗ	Н	н
XA735	снз-	H <sub>3</sub> C H <sub>3</sub> C-{}-{	Н	СНЗ	н	Н
XA736	СНЗ	H <sub>3</sub> C	н	снз	н	н
XA737	снэ-	F_F	Н	СНЗ	н	н
XA738	снз-	F√∰	н	снз	н	н
XA739	снз–	F F	н	СНЗ	н	н
XA740	снз–	F	Н	снз	н	н
XA741	снз-	F.	Н	СНЗ	н	н
XA742	CH3-	F +	н	СНЗ	н .	н
XA743	CH3-	CI_CI	н	СНЗ	н	H
XA744	CH3-	GH CH	Н	<b>СН3</b>	Н	н



### PCT/JP2004/004320

No.	R1	R2	R3	R4	R5	R6
XA745	снз–	a a	н		н	Н
XA746	CH3-	G G a	Н	СНЗ	Н	Н
XA747	снз–	a, ci—⟨	н	снз	<b>H</b>	Н
XA748	СН3	a G	н	СНЗ	Н	Н
XA749	CH3-	H <sub>3</sub> CO_OCH <sub>3</sub>	Н	снз	Н	H
XA750	СН3-		н	снз	Н	Н
XA751	СН3-	H²∞ → H²∞	н	снз	Н	Н

	-					
No.	R1	R2	R3	R4	R5	R6
XA752	СН3-	OCH <sub>3</sub>	н	снз	Н	Н
XA753	СН3-	H <sub>3</sub> CO	н	СНЗ	н	H
XA754	СН3	H <sub>3</sub> CO	н	СНЗ	н	н
XA755	снз-	F_OCH <sub>3</sub>	н	СНЗ	н	H
XA756	снз-	OCH <sub>3</sub> F—⟨◯)→	н	СНЗ	н	Н
XA757	СН3	OCH₃ F-{\rightarrow}-{\righta	н	СНЗ	н	н
XA758	СН3-	OCH <sub>3</sub>	н	снз	н	Н
XA759	CH3-	OCH <sub>3</sub> F	н	СНЗ	н	Н
XA760	снз-	OCH₃ F	н	СНЗ	н	Н
XA761	CH3-	H₃CQ F—	н	СНЗ	н	Н
XA762	СН3-	H₃CQ F	Н	снз	н	н
XA763	CH3-	H₃CO_F	н	СНЗ	н	н
XA764	СН3-	H₃CO-⟨¯¯⟩}	н	СНЗ	Н	Н
XA765	СН3-	н₃∞	H	СНЗ	Н	н
XA766	снз–	H₃CO-{}}	н	снз	н	н

•	· · · · · · · · · · · · · · · · · · ·	1			Ta.	
No.	R1	R2	R3	R4 .	R5	R6
XA767	снз-	CI_OCH <sub>3</sub>	н	снз	Н	н
XA768	СН3-	OCH <sub>3</sub>	н	снз	Н	Н
XA769	CH3-	α σαн³	н	снз	Н	Н
XA770	СН3-	CI CCH³	H	снз	н	Н
XA771	снз-	H₃CQ CI—⟨¯)—;	н	снз	Н	Н
XA772	CH3-	H₃CO CI	Н	снз	Н	Н
XA773	СН3-	H₃CO_CI	н	снз	н	н

No.	R1	R2	R3	R4	R5	R6
XA774	снз-	CI H₃CO-⟨¯)—}	Н	снз	н	н
XA775	СН3-	CI → H₃co	н	СНЗ	н	н
XA776	СН3-	CI H₃CO-⟨¯_}~;	н	снз	н	Н
XA777	СН3-	F_CH <sub>3</sub>	Н	снз	H ·	Н
XA778	СН3-	CH <sub>3</sub> F—{	н .	снз	н	Н
XA779	СН3-	CH <sub>3</sub> F	Н	снз	н	Н
XA780	СН3-	CH₃ C F	Н	СНЗ	н	Н
XA781	СН3-	H <sub>3</sub> C F—{}}	Н	снз	н	Н
XA782	СН3	H₃C	Н	СНЗ	H	н
XA783	CH3	H <sub>3</sub> C F	н	снз	Н	Н
XA784	СН3-	H₃C-⟨S}}	Н	снз	Н	Н
XA785	снз-	H <sub>3</sub> C	Н	снз	н	н
XA786	СН3-		Н	снз	Н	н
XA787	СН3-	Br_OCH <sub>3</sub>	Н	снз	Н	н
XA788	С <b>Н3</b> -	OCH₃ Br—{}	Н	снз	Н	Н



# PCT/JP2004/004320

No.	R1	R2	R3	R4	R5	R6
XA789	CH3-	OCH <sub>3</sub> Br	н	СНЗ	н	н
XA790	снз-	OCH <sub>3</sub>	Н	СНЗ	н	Н
XA791	снз-	H <sub>3</sub> CO Br—{	Н	СНЗ	Н	н
XA792	СН3-	H₃CO Br	Н	СНЗ	н	Н
XA793	снз–	H <sub>3</sub> CO_Br	н	СНЗ	Н	Н
XA794	CH3-	H₃CO-{\bigset}	н	СН3	н	н
XA795	CH3	H <sub>3</sub> CO	н .	СН3	н	Н

No.	R1	R2	R3	R4	R5	R6
110.	.,,	Bς	140	114	110	
XA796	снз-		H	СНЗ	н	н
XA797	СН3-	H³CO }	н	СНЗ	Н	н
XA798	CH3-	Cn-{_}}- Cu-{_}}-	н	СНЗ	Н	н
XA799	CH3-	CN-⟨_}OCH3	Н	СНЗ	н	н
XA800	CH3-	H³CO }	н	СНЗ	Н	Н
XA801	снз-	H <sub>3</sub> CO	н	СНЗ	н	Н
XA802	CH3-	CN → OCH3	Н	СНЗ	н	Н
XA803	CH3-	F-CS-I F	н	СНЗ	н	Н
XA804	снз-	OCH <sub>3</sub> F-{_}	н	СНЗ	Н	Н
XA805	CH3-	H₃CO-{_}_}F F	н	снз	Н	н
XA806	СН3	OCH <sub>3</sub> F-⟨_}; OCH <sub>3</sub>	Н	снз	н	Н
XA807	СН3	OCH3 OCH3	Н	СНЗ	н	н
808AX	СН3-	a- <a>Ca&gt; a</a>	н	CH3	Н	Н
XA809	снз-	осн₃ cı—(i cı	н	СНЗ	Н	н
XA810	СН3-	а н₃∞-⟨∑-; a	Н	снз	н	н

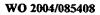
No.	R1	R2	R3	R4	R5	R6
XA811	СН3-	OCH³	н		н	Н
XA812	СН3-	OCH3 H³CO-⟨¯_)→1 OCH3	н	снз	Н	н
XA813	CH3		н	снз	н	Н
XA814	СН3-	H <sub>3</sub> CO	н	СНЗ	Н	н
XA815	CH3-	H3CO-{\}-{\}-{\}-{\}-	н	снз	Н	н
XA816	снз-	OCH <sub>3</sub> }	Н	снз	Н	Н
XA817	СН3-	H <sub>3</sub> CQ ,	Н	снз	H	н

No.	R1	R2	R3	R4	R5	R6
XA818	снз-	H <sub>3</sub> CO-{\}_\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н			Н
XA819	СН3-	OCH <sub>3</sub>	н	СНЗ	н	Н
XA820	снз-	H,00	н	СНЗ	H	н
XA821	СН3-	H₃CO-{_}	Н	снз	Н	Н
XA822	СН3-	<b>₫</b>	н	снз	H	Н
XA823	снз-		н	снз	H	н
, XA824	СН3-	F-()-{)-{	Н	снз	Н	Н
XA825	снз-		н	снз	н	Н
XA826	снз-		Н	снз	н	н
XA827	снз–		H	снз	Н	н
XA828	снз-	Q-0	Н	СНЗ	Н	н
XA829	СН3-	\$\_\O	н	снз	Н	н
XA830	СН3-		Н	СНЗ	Н	н
XA831	СН3~	Q ·	н	СНЗ	н	Н
XA832	СН3-		Н	СНЗ	н	Н



No.	R1	R2	R3	R4	R5	R6
XA833	снз-		H	снз	Н	н
XA834	CH3-		н	СНЗ	н	н
XA835	снз-		н	СНЗ	Н	Н
XA836	СН3-	T)	Н	снз	н	н
XA837	СН3-		н	снз	н	Н
XA838	СН3-		Н	снз	Н	Н
XA839	СН3-		н	СНЗ	Н	Н

No.	R1	R2	R3	R4	R5	R6
XA840	СН3		н	СНЗ	н	н
XA841	снз-	Co	Н	СН3	н	н
XA842	снз-	(C)	Н	СНЗ	Н	н
XA843	снз-	,CC	н	СНЗ	н	н
XA844	снз-	Ĉ.	н	СНЗ	н	н
XA845	снз-		н	СНЗ	н	Н
XA846	снз-		Н	СНЗ	н	н
XA847	снз-		Н	СНЗ	н	н
XA848	снз–	(CI)	Н	СНЗ	н	н
XA849	снз-	,CIŞ	н	СНЗ	н .	н
XA850	снз–	Ţ\$	Н	СНЗ	Н	н
XA851	снз–		н	СНЗ	Н	н
XA852	снз-		н	СНЗ	н .	н
XA853	СН3-	T	н	снз	Н	н
XA854	снз-		н	снз	н	н
XA855	снз-	ŢŢ,	н	снз	н	н
XA856	снз-	(Try)	н	снз	н	н
XA857	снз-		н	СНЗ	Н .	н
XA858	снз	T N	Н	снз	Н	н
XA859	снз-	CT)+	Н	снз	н	н
XA860	снз-	J.	Н	СНЗ	н	н





	D4:		<b>D</b> 0	D4	DE	ne
No.	R1	R2	R3	R4	R5	R6
XA861	снз-	<b>(C)</b>	н	СНЗ	н	н
XA862	снз-	, Ci	н	СНЗ	н	н
XA863	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СНЗ	н	н
XA864	снз–	O'S'	н	сня	н	н
XA865	С <b>Н3</b> –	Ţ, N,	H	СНЗ	н	н
XA866	снз–	TOS.	Н	СНЗ	н	Н
XA867	снз–	'\Q\\	Н	СНЗ	н	Н
XA868	снз-	QT'S	н	СНЗ	н	Н
XA869	снз-	C;	н	снз	Н	H
XA870	снз-	Ţ,	н	снз	н	H
XA871	снз–	TO:	Н	снз	н	н
XA872	снз–	,CT	Н	снз	н	н
XA873	снз–	Ĩ,	Н	снз	н	Н
XA874 ·	снз–	OZ)	н	снз	Н	Н
XA875	СН3-	Č;	Н	снз	н	н
XA876	СН3-	. CP	н	снз	Н	н
XA877	СН3	,CT3 <sup>1</sup>	н	снз	Н	н
XA878	снз-	Ť,	н	снз	н	н
XA879	снз–	Č.	н	снз	н	Н
XA880	снз-	'CS	н	СНЗ	н	н
XA881	снз-	TOR	н	снз	Н	н
XA882	СН3-	Č;	н	снз	н	н

No.	RI	R2	R3	R4	R5	R6
XA883	снз-	CH3-	Н		н	<b>н</b>
XA884	снз-	снзсн2-	Н		н	н
XA885	CH3-	<b>~</b>	н		н	н
XA886	снз-	$\nearrow$	Н		Н	н
XA887	снз-	<b>~</b> ~	Н		н	Н
XA888	снз-	人、	Н		н	н
XA889	снз-	$\gamma$	н		Н	н
XA890	снз-	丫	Н		н	н
XA891	CH3-	<b>^</b>	н	Qu	н	н
XA892	снз-	<b>**</b>	н	Q	Н	н
XA893	снз-	\\ \ .	н	Qu	Н	н
XA894	снз-	7	н	Qu	н	н
XA895	снз-	~~~	н	Qu	н .	н
XA896	снэ-	<del></del>	н		н	н
XA897	СН3-	~~~ <u>`</u>	н		н	н
XA898	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Qu	н	Н
XA899	снз-	n-C8H17-	н .	Qu	н	Н
XA900	снз-	L	н	Qu	н	н
XA901	снз-	Qu	н		н	н
XA902	снз-		н	Qu	н	н
XA903	снз-	0~~	Н		н	н
XA904	СН3-	D-1	н	Qu	н	н

No.	RI	R2	R3	R4	R5	R6
XA905	снз-		Н		Н	н
XA906	снз-	$\bigcirc$	н	Q	Н	н
XA907	снз-	$\bigcirc$ $\dashv$	н	Qu	Н	н
XA908	СН3-	$\bigcirc$	н		н	Н
XA909	снз–		н	Q	Н	н
XA910	СН3-		н	Q	Н	н
XA911	снз–	<b></b>	н	Qu	н	н
XA912	снз-		н	Q	н	н
XA913	снз-		н	Q	н	н
XA914	снз–	F-(	н	Q	н	н
XA915	CH3-		н	Q	Н	н
XA916	снз-	F-On-4	н	Qu	н	Н
XA917	СН3-	CI	н	Qu	н	Н
XA918	снз-	CI C)	н	Qu	н .	Н
XA919	СН3-	c⊢ <u>(</u>	H	Q	н	н
XA920	снз–	C⊢ <b>(</b> )~-1	н	Qu	Н	н
XA921	СН3-	C⊢{\right\}{	н	Qi	н	н
XA922	снз–	Br	Н	Qu	Н	н
XA923	снз-	Br.	н	Qu	Н	н
XA924	снз-	Br-{}-{	Н	Qu	Н	н
XA925	снз-	Br-{}-{	н	Qu	н	н
XA926	СН3	Br— <b>(_</b> )⊪{	Н	Q	Н	н

No.	RI	R2	R3	R4	R5	R6
		,!				
XA927	CH3-		Н		Н	Н
XA928	снз-		н	Qu	н	н
XA929	снз-		н	Qu	н	н
XA930	снз-	СН₃	н	Q	н	н
XA931	CH3-	H₃C	н	Q	н	H
XA932	снз-	H₃C- <b>(_</b> )—{	н		Н	н
XA933	снз-	C <sub>2</sub> H <sub>5</sub> -{}-{	н	Q	н	Н
XA934	CH3-	n-C₃H <sub>7</sub> -∕€}{	н	Q	Н	Н
XA935	снз-	n-C <sub>4</sub> H <sub>9</sub> -	Н	Q	н	н
XA936	СН3	OH OH	Н		н	Н
XA937	СН3-	HO ————————————————————————————————————	н		н	н
XA938	снз-	HO-{\right\}-{	н	Q	н	Н
XA939	снз-	OCH₃	н	Qu	н	Н
XA940	снз-	H₃CO ————————————————————————————————————	н		н	Н
XA941	снз-	H³CO-{_}-{	н	Qu	н	н
XA942	снз-	H₃CO- <b>{</b> }-{	н	Q	н	н
XA943	снз-	H <sub>3</sub> CO-{\rightarrow}m{\frac{1}{2}}m{\frac{1}{2}}	н		н	н
XA944	CH3-	OC₂H <sub>5</sub>	Н	Qu	н	н
XA945	СН3-		Н	Qu	н	н .
XA946	снз-	C <sub>2</sub> H <sub>5</sub> O-{}-{	Н		н	Н
XA947	СН3-	n-C₃H <sub>7</sub> O- <b>⟨</b> }—{	н		н	н
XA948	СН3-	n-C <sub>4</sub> H <sub>9</sub> O-	н	Qu	н	н

XA949       CH3-       NO2	No.	R1	R2	R3	R4	R5	R6
XA950 CH3	XA949	снз-	\_/ <u>`</u>	н	Qu	н	Н
XA952       CH3-       CN       H	XA950	снз-	O <sub>2</sub> N	н	Qu	н	н
XA952       CH3-       CH3-       H <t< td=""><td>XA951</td><td>снз-</td><td></td><td>н</td><td>Qu</td><td>Н</td><td>H</td></t<>	XA951	снз-		н	Qu	Н	H
XA953       CH3-       CH3-       H <t< td=""><td>XA952</td><td>снз-</td><td></td><td>н</td><td>Qu</td><td>н</td><td>н</td></t<>	XA952	снз-		н	Qu	н	н
XA955 CH3-	XA953	снз-	NC	н	Qu	н	H
XA955       CH3-       GH3-       F3C       H	XA954	снз-		н	Qu	н	н
XA956       CH3-       F3C-       H <t< td=""><td>XA955</td><td>снз-</td><td></td><td>Н</td><td>Qu</td><td>н</td><td>Н</td></t<>	XA955	снз-		Н	Qu	н	Н
XA958       CH3-       COOH       H <t< td=""><td>XA956</td><td>снз</td><td>F<sub>3</sub>C</td><td>н .</td><td>Qu</td><td>н</td><td>н .</td></t<>	XA956	снз	F <sub>3</sub> C	н .	Qu	н	н .
XA958       CH3-       CH3-       H <t< td=""><td>XA957</td><td>СН3-</td><td></td><td>н</td><td>Qu</td><td>н</td><td>н</td></t<>	XA957	СН3-		н	Qu	н	н
XA969       CH3-       H	XA958	CH3-	<b>◯</b> -₁	н	Qu	н	н
XA961       CH3-       CO₂Me       H       <	XA959	снз-	HOOC	н	Qu	н	н
XA961       CH3-       H	XA960	снз-		н		н	Н
XA962       CH3-       MeO₂C-       H	XA961	СН3-		н	Qu	Н	н
XA964       CH3-       CO₂Et       H       <	XA962	снз-	MeO <sub>z</sub> C	н	Q	Н	н
XA964       CH3-       CH3-       H <t< td=""><td>XA963</td><td>СН3-</td><td></td><td>Н</td><td>Q</td><td>Н</td><td>н</td></t<>	XA963	СН3-		Н	Q	Н	н
XA965 CH3-	XA964	СН3-	<b>⟨</b> }~₁	н .		н	H
XA967 CH3- SMe H H H  XA968 CH3- MeS H H H H  XA969 CH3- MeS- H H H H	XA965	СН3-		Н	Qu	H	Н
XA967 CH3- H H H  XA968 CH3- MeS H H H H  XA969 CH3- MeS- H H H H	XA966	СН3-		Н	Q	н	н
XA968 CH3- H H H  XA969 CH3- MeS- H H H  SO <sub>2</sub> Me	XA967	СН3-		Н	Qi	н	н
SO <sub>2</sub> Me	XA968	снз-	MeS	Н	Q	н	н
хаэ70 CH3- SO <sub>2</sub> Me H H	XA969	снз-		Н	Qu	н	н
	XA970	снз-	SO₂Me	н	Qu	н	н



No.	R1	R2	R3	R4	R5	R6
110.		MeO <sub>2</sub> S	<del>                                     </del>			
XA971	снз-		Н		н	н
XA972	снз–	MeO₂S-{_}	Н	Or	н	н
XA973	снз-	NH <sub>2</sub>	н .		н	н
XA974	снз-	H <sub>2</sub> N	н	Qr	н	Н
XA975	снз-	H <sub>2</sub> N-{}-{	Н		Н	Н
XA976	снз-	NMe <sub>2</sub>	н		н	н .
XA977	снз-	Me <sub>2</sub> N	Н	Q	н	н
XA978	CH3-	Me <sub>2</sub> N-()	н	Qu	н	н
XA979	снз-		Н	Qu	н	Н
XA980	снз-	(n-()	н		н	н
XA981	снз-	CN-C)-1	н	Q	н	н
XA982	снз-		н		н	н
XA983	снз-		Н		н	н
XA984	СН3-		н		н	н
XA985	снз-		н		н	н
XA986	снз-	O+Q	н	Q	н	н
XA987	снз-	<b>€</b> ₩€	н		н <sub>.</sub>	н
XA988	снз-	H3CN_N-	н	Qu	н	Н
XA989	снз-	H <sub>3</sub> CN N-	н	Qu	н	н
XA990	снз-	H³CN N-{}	н	Q	н	н
XA991	снз-	H <sub>3</sub> C CH <sub>3</sub>	Н	Q	н	н
XA992	снз-	H <sub>3</sub> C-{}_{CH <sub>3</sub>	н	Q	н	н

No.	R1	R2	R3	R4	R5	R6
	снз-	CH₃ → H₃C	Н	Ö.	Н	Н
XA994	снз-	CH₃	н	Q	Н	н
XA995	СН3-		Н		Н	н
XA996	СН3-	H <sub>3</sub> C H <sub>3</sub> C	Н		н	Н
XA997	СН3-	F-F	н		н	н
866VX	СН3-	F——F	н		Н	Н
XA999	CH3-	ŞĞ-	Н	Q	н	н
XA1000	снз-	ÇF; F	Н	Q	н	н
XA1001	CH3-	F—————————————————————————————————————	н		Н	н
XA1002	снз-	F F	н		Н	н
XA1003	снз-	CI_CI	н		Н	H
XA1004	CH3-	CI—(CI	н		н	н

-		The	Tea		155	1==
No.	Rt	R2	R3	R4	R5	R6
XA1005	снз-	a CI	н		н	н
XA1006	снз-	CI CI	н	Q	н	н
XA1007	снз-	CI-	Н	Q	н	н
XA1008	снз-	CI CI	н	Q	н	Н
XA1009	снз-	H <sub>3</sub> CO_OCH <sub>3</sub>	н	Q	<i>.</i> н	н
XA1010	СН3-	осн <sub>3</sub> н₃со-{∑}-⊰	Н	Q	н	Н
XA1011	CH3-	OCH₃ H₃CO	н	Q	н	н
XA1012	CH3-	OCH <sub>3</sub>	н	Q	Н	н
XA1013	СН3-	H₃CO-⟨¯}-⊀	н	Q	Н	н
XA1014	СН3-	H <sub>3</sub> CO	Н	Q	Н	н

No.	ID1	Im	Ina	154	T	
NO.	R1	R2	R3	R4	R5	R6
XA1015	снз-	F_OCH <sub>3</sub>	н	Q	н	Н
XA1016	снз-	OCH <sub>3</sub>	н	Q	Н	н
XA1017	СН3-	OCH <sub>3</sub>	н		н	н
XA1018	СН3-	OCH <sub>3</sub>	н	Q	н	Н
XA1019	СН3-	OCH <sub>3</sub>	н	Q.	н	н
XA1020	снз-	OCH <sub>3</sub>	H	Q	н	н
XA1021	снз-	H₃CO F—√	Н	Q	н ,	н
XA1022	снз-	H₃CO F	н	Q	н	Н
XA1023	СН3	H <sub>3</sub> CO_F	н	Q	н	н
XA1024	СН3-	H₃CO-{\bigs_}	Н	Q	Н	н
XA1025	снз–	F → H₃co	H		H	Н
XA1026	CH3-	H₃CO-{}}	Н		H	н

No.	R1	R2	R3	R4	R5	Inc
XA1027	СН3-	CI_OCH <sub>3</sub>	Н	Ö.	Н	R6 H
XA1028	СН3-	CI—⟨SH3	н	Q	н	н
XA1029	СН3-	OCH <sub>3</sub>	н	Q	н	н
XA1030	снз-	OCH <sub>3</sub>	н	Q	н	н
XA1031	снз-	H <sub>3</sub> CO CI—	н	Q	н	н
XA1032	СН3-	H₃CO CI	н	Q	н	Н
XA1033	СН3	H₃CO_CI	Н	Q	н	н
XA1034	СН3-	H₃CO-⟨∑)—{	н	Q	H	Н
XA1035	CH3-	CI H <sub>3</sub> CO	Н	Q	н	Н
XA1036	СН3~	CI H₃CO-⟨¯¯ <mark>&gt;</mark> →{	Н	Q	н	Н

No.	R1	R2	R3	R4	R5	R6
XA1037	СН3-	F_CH <sub>3</sub>	Н		Н	Н
XA1038	СН3-	CH <sub>3</sub> F—{}	н		Н	н
XA1039	снз-	CH₃	н	Q	н	н
XA1040	СН3-	CH <sub>3</sub> ← F	н	Q	н	Н
XA1041	снэ-	H <sub>3</sub> C F-\	н	Q	Н	Н
XA1042	снэ–	H <sub>3</sub> C ↓ F	Н	Q	н	н
XA1043	СН3	H <sub>3</sub> C_F ←	Н	Q	н	н
XA1044	снз-	H₃C-⟨S	н	Q	н	Н
XA1045	СН3-	H₃C F	Н	Q	Н	н
XA1046	снз-	F_ H₃C-	н	Q	Н	н
XA1047	снз–	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Q	Н	- Н
XA1048	снз-	OCH₃ Br—€	н		Н	Ĥ



No.	R1	R2	R3	R4	R5	R6
XA1049	снз-	OCH <sub>3</sub> Br	н	Q	Н	Н
XA1050	снз-	OCH <sub>3</sub> ⇒ Br	Н		н	Н
XA1051	сн <b>з</b> -	H₃CQ Br—⟨¯¯¯)—;	Н		н	Н
XA1052	снз-	H₃CO Br	Н		Н	н
XA1053	снз	H <sub>3</sub> CO_Br	Н		Н	н
XA1054	снэ-		Н	Q	н	н
XA1055	CH3-	Br √-} H₃co	н	Q	н	н
XA1056	СН3	H <sub>3</sub> CO	н	Q	Н	H
XA1057	снз-	H <sub>3</sub> CO	н		Н	Н
XA1058	СН3	OCH <sub>3</sub>	Н .		Н	Н



No.	R1	R2	R3	In.	Ins	
XA1059	СН3-	CN-C>-och		R4	R5 H	R6 H
XA1060	СН3-	H <sub>3</sub> CO >	Н	Q	н	н
XA1061	снз–	H <sub>3</sub> CO	н	Qu	н	н
XA1062	СН3-	OCH3	н		н	Н
XA1063	снз-	F—————————————————————————————————————	H .	Q	н	Н
XA1064	снз-	OCH <sub>3</sub> F————————————————————————————————————	Н	Q	Н	Н
XA1065	СНЗ-	H₃CO-(SH) F	H	Q	Н	н
XA1066	снз–	OCH <sub>3</sub>	Н	Q	н	н
XA1067	снз-	OCH3	H		Н	Н
XA1068	снэ–	CI	Н		Н	Н
XA1069	СН3-	CI.	н	Q	Н	Н
XA1070	СН3-	H₃co-{_}; a	Н	Q	н	Н

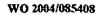


No.	R1	R2	R3	R4	R5	R6
XA1071	снз-	OCH <sub>3</sub> CH	н	Q	Н	Н
XA1072	снз-	OCH <sub>3</sub>	н	Q	Н	н
XA1073	снз–	OCH <sub>3</sub>	н	Q	Н	н
XA1074	сн3-	H <sub>3</sub> CQ	н	Q	н	н
XA1075	СН3-	н₃со-{`}-{`}-∤	н		н	Н
XA1076	снз-	OCH <sub>3</sub> }	н	Q	Н	н
XA1077	CH3-	H <sub>3</sub> CO	Н	Qr	Н	н
XA1078	CH3-	н₃∞-(∑-(С)	Н	Q	н	Н
XA1079	CH3-	OCH <sub>3</sub>	Н	Q	Н	н
XA1080	снз-	H <sub>3</sub> CQ	н	Q	H	H.

No.	R1	IR2	R3	R4	R5	R6
140.	<u> </u>	63 63	1/13	In4	149	NO
XA1081	CH3-	H3CO-(_)-(/_)	н	Qr_	н	н .
XA1082	снз-	₫-0-1	н	Q	н	н
XA1083	СН3-		н	Q	Н	н
XA1084	снз-	F-()-()-1	Н	Q	Н	н
XA1085	CH3-	<u>a</u> -a	Н	Q	н	н
XA1086	снз-	<b>5</b> -0	н	Q	н	н
XA1087	СН3-	F-O-O'	н	Q	Н	н
XA1088	снз-	Q.O	н	Q	н	н
XA1089	снз	ρΏ	н	Q	Н	н
XA1090	снз-		н	Q	н	н
XA1091	снз-		н .	Q	Н	н
XA1092	снз-	CC,	н	Q	н	н
XA1093	СН3-		Н	Qu	Н	н
XA1094	СН3		н	Q	н	н
XA1095	СН3	Č,	н	Q	н	н
XA1096	снз–	O	н	Q	н	H·
XA1097	СН3-	,O)	н	Qu	н	н
XA1098	снз-	Ŷ	н	Qi	Н	Н
XA1099	снз–		н	Q.	Н	н
XA1100	снз–	CI)	н	Qu	Н	н
XA1101	снз-	Ċ;	н	Q	н .	н
XA1102	снз-	TO:	н		Н	н



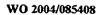
No.	R1	R2	R3	R4	R5	R6
XA1103	CH3-	,CC),	н	Q	н	н
XA1104	СН3-	ÇI;	н	Q	н	н
XA1105	снз-	CI\$+	н	Q	Н	н
XA1106	снз-	O.	н	Q	н	н
XA1107	СН3		н	Q	н.	- H
XA1108	снз-	TI	н		н	н
XA1109	снз-	,CI3	н		н .	. н
XA1110	СН3-	Ĉ.	н		н	н
XA1111	снз-	C.	н	Q.,	H	н
XA1112	СН3-	<b>Ö</b>	н		н	н
XA1113	СН3-	TOP	н		н	н
XA1114	снз-	, CY	н	Qi	н	Н.
XA1115	снз-	ŢŢ,	н		Н	н
XA1116	CH3-		н		н .	н
XA1117	CH3		н		- н	н
XA1118	CH3-	TON,	Н	Q	Н	н
XA1119	СН3-		н	Qu	н	н
XA1120	CH3-	Ž,	н	Qu	н	н
XA1121	снз-	TOP	н		Н	н
XA1122	снз-	, CP3	Н	Qu	н	н
XA1123	снз-		Н	Qu	н	н
XA1124	CH3-	(Is)	н		н	н





No.	R1	R2	R3	R4	R5	R6
XA1125	СН3-	J.N.	н	Q	Н	н
XA1126	СН3-	'O's	н	Q	н	н
XA1127	снз-	, II's	Н		н	Н
XA1128	СН3-	Ţ <sup>s</sup>	Н	Qu	н	н
XA1129	CH3-	OF)	Н	Qu	н	H.
XA1130	снз-		н	Qu	н	н
XA1131	СН3-		н	Qu	н	н
XA1132	CH3-	,CC;	н .	Qu	н	н
XA1133	снз-	Ğ.	Н	Qu	н	н
XA1134	снз-	O.	н	Qi	н	Н
XA1135	СН3-		н	Qi	н	н
XA1136	снз-	TO Y	Н	Qi	н	н
XA1137	CH3-	,CT3"	н	Qi	н	н
XA1138	CH3-	ČL3,	н	Q.	н	н
XA1139	CH3-	Č.	Н	Q	Н	н
XA1140	CH3-	,CC	Н	Q	н	н
XA1141	снз-	CO	н		н	н
XA1142	снз-	Ğ;	Н		н	н
XA1143	снз–	CH3-	н	Ŷ,	н	н
XA1144	СН3-	снзсн2-	н	2,	Н	н
XA1145	СН3	~\`\	н	Ŷ,	Н	н
XA1146	СН3-	Y	н	Ŷ,	н	н

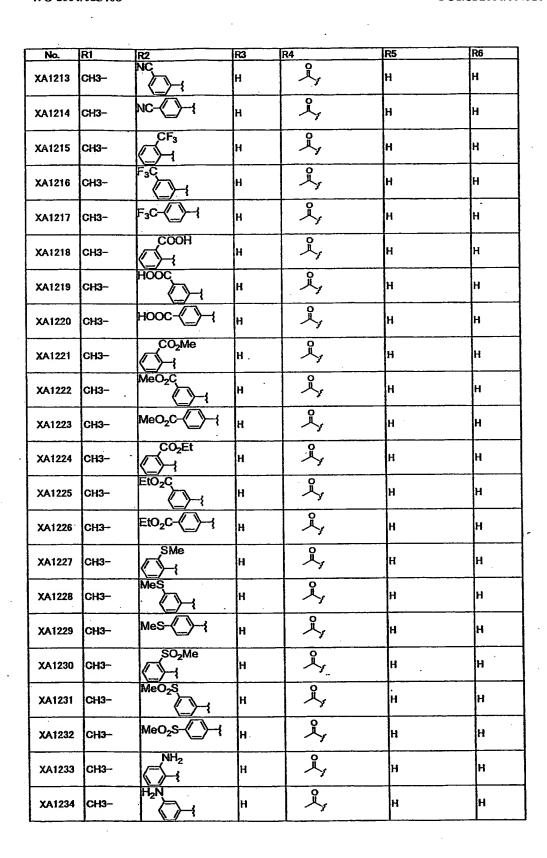
No.	R1	R2	R3	R4	R5	R6
XA1147	снз-	<b>\\\\</b>	н	Ŷ,	н	н
XA1148	СН3	Į,	н	Ů,	н	н
XA1149	снз-		Н	Ŷ,	н	н
XA1150	снз-	Y	Н	Ŷ,	н	н
XA1151	снз-	~~\\	н	Ŷ,	н	H
XA1152	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	2	н	Н
XA1153	снз-	Xv.	Н	Ŷ,	н	н
XA1154	снэ–	7	н	Î,	н	н .
XA1155	СН3-	~~~``	H	Ŷ,	н	Н
XA1156	сна-	L~~	Н	Ŷ,	н .	н
XA1157	снэ-	~~~``	н	l,	н	н
XA1158	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	ي.	H	н
XA1159	СН3-	n-C8H17-	н	Ŷ,	н	н
XA1160	СН3-	L~~~	н	<u></u>	н	н
XA1161	снз-		н	Ly	н	н
XA1162	СН3-		н	Ŷ,	н	н
XA1163	снз-		н	Ŷ,	н	н
XA1164	СН3-	<b>→</b>	н	گ <sub>ا</sub>	н	н
XA1165	СН3-	$\Diamond$	Н	· Å,	н	н
XA1166	СН3-	$\bigcirc$	Н	2,	Н	н
XA1167	снз-	$\bigcirc$ $\dashv$	Н	Ŷ,	Н	н
XA1168	CH3-	$\bigcirc$ $\vdash$	н	Î,	н	н





No.	R1	R2	R3	R4	R5	R6
XA1169	снз-		Н	Ļ,	н	н
XA1170	снз-		Н	Ŷ,	н	н
XA1171	снз-		н	Ŷ,	Н	н
XA1172	снз-		н	ي ا	н	н
XA1173	снз-		н	گ <sub>ه</sub>	н	н
XA1174	снз-	F-()-1	н	گ <sub>ه</sub>	н	н
XA1175	снз-		H	گہ	н	н
XA1176	СН3-	F-{\_}\\\	н	Ŷ,	н	н
XA1177	снз-	CI	н	Ŷ,	Н	н
XA1178	снз-	CI CI	н	Ŷ,	н	н.
XA1179	СН3-	C⊢ <b>(</b> ){	н	<sup>گ</sup> ہ	н	н
XA1180	снз-	c⊢ <b>(</b> )→1	н	Ŷ,	Н	н
XA1181	снз-	CI—(	н	بُ	н	н
XA1182	СН3-	Br —{	H	Ŷ,	Н .	н
XA1183	снз-	Br.	н	Ŷ,	н	н
XA1184	снз-	Br-{}-{	н	ŝ,	Н	н
XA1185	снз	Br-C>-{	н	l,	Н	н
XA1186	снз-	Br—{	н	Ŷ,	н	н
XA1187	снз-		н	<u>گ</u>	н	н
XA1188	снз-		н	Ž,	Н	н
XA1189	снз-		Н	<u></u>	Н	Н
XA1190	снз-	СН₃ СУ⊣∤	н	ئى ا	Н	н

No.	R1	Ipa	R3	R4	R5	R6
140.	KI	R2 H₃C			160	No
XA1191	снз-		н	<b>1</b> ,	н	н
XA1192	снз-	H <sub>3</sub> C-{}-{	н	<b>3</b> ,	н	Н
XA1193	снз-	C <sub>2</sub> H <sub>5</sub> -{}-{	н	Ŝ,	н	н
XA1194	снз-	n-C <sub>3</sub> H <sub>7</sub> -{	н	i,	н	н
XA1195	СН3-	n-C <sub>4</sub> H <sub>9</sub> -{}-{	н	Ŷ,	н	н
XA1196	снз-	OH ⟨=⟩→	н	گہ	H	н
XA1197	снз-	HO	н	گړ.	н .	Н
XA1198	снз-	HO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar	Н	Ŷ,	н	н
XA1199	СН3-	OCH₃	н	Ŷ,	н	н
XA1200	снз-	H₃CQ 	н	ڳ <sub>ي</sub>	Н	н
XA1201	CH3-	H₃CO- <b>⟨</b> }-{	н	گ <sub>ا</sub>	н	н
XA1202	снз-	H3CO-{>-1	н	Å,	н	н
XA1203	CH3-	H3CO-{\bigs\}\!\!\!	н	Ŷ,	н	н
XA1204	снз-	OC <sub>2</sub> H <sub>5</sub>	н	Ŝ,	н	н
XA1205	СН3	C₂H₅O ⟨}-{	н	Ŷ,	н	н
XA1206	CH3-	C2H2O-{}-{	н	3,	н	Н
XA1207	снз-	n-C <sub>3</sub> H <sub>7</sub> O-	н	Ŷ,	н	н
XA1208	СН3-	n-C <sub>4</sub> H <sub>9</sub> O-	н	Ŝ,	н	н
XA1209	CH3-	NO <sub>2</sub>	Н	Î,	н	н
XA1210	снз-	O <sub>2</sub> N	н .	ů,	н	н
XA1211	снз-	02N-()-1	н	Ŷ,	н	н
XA1212	СН3-	CN	н	Î,	н	н



No.	R1	R2	R3	R4	R5	R6
XA1235	СН3-		H	Ŷ,	н	Н
XA1236	снз-	NMe₂ →	H	Î,	Н	Н
XA1237	снз-	Me <sub>2</sub> N	Н	Ŷ,	н	н
XA1238	снз-	Me <sub>2</sub> N-{	Н	l,	н	Н
XA1239	СН3-		н	l,	Н	Н
XA1240	СН3-		н	گي	н	Н
XA1241	снз-		н	Ŷ,	Н	н
XA1242	CH3-		н	L,	н	Н
XA1243	СН3-		н	l,	н	н
XA1244	СН3-		Н	<u>گ</u>	н	Н
XA1245	СН3-		н	١,	H	Н
XA1246	снз-	<->	н	l,	н	н
XA1247	снз-	<b>€</b> N-€}-1	н	گي	Н	Н
XA1248	CH3	H3CN_N-	н	l,	Н	Н
XA1249	. СН3-	H3CN N-	, н	٤,	н .	н

Н

Н

н

н

XA1255 CH3-

XA1256 CH3-

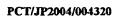
No.	R1	R2	R3	R4	R5	R6
XA1250	снз-	H₃CN N-(S)-1	Н	<u></u>	н	н
XA1251	снз-	H₃C_CH₃	н	l,	н	н
XA1252	СН3-	H <sub>3</sub> C-{\rightarrow}-\footnote{\rightarrow}-\footnote{\rightarrow}	Н	١,	н	н
XA1253	снз-	CH <sub>3</sub> H <sub>3</sub> C	Н	2,	н	Н
XA1254	CH3-	CH <sub>3</sub>	н	l,	н	н

Ŷ,

н

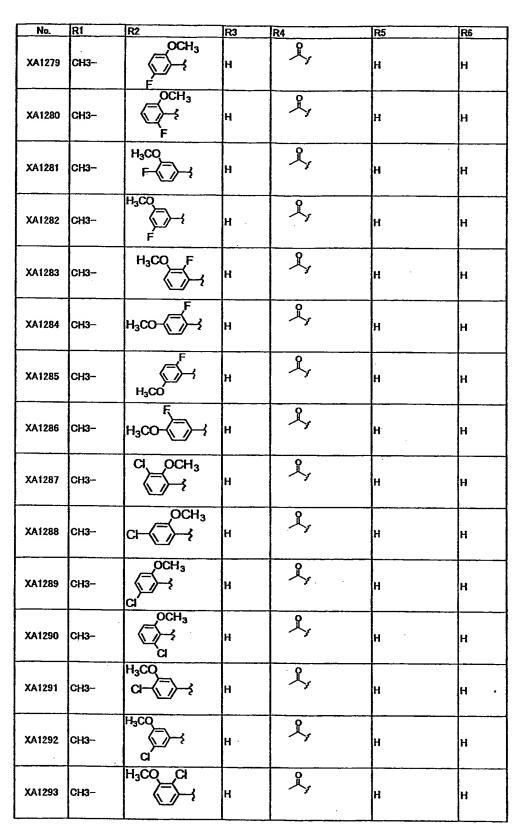
No.	R1	R2	R3	R4	R5	R6
XA1257	СН3-	F_F	Н	Ŷ,	н	Н
XA1258	снз-	F————	Н	Ŷ,	Н	н
XA1259	снз-	F.	н	Ŷ,	н	н
XA1260	СН3-	F F	Н	Ŷ,	Н	н
XA1261	CH3-	F.	н	Ŷ,	н	н
XA1262	СН3-	F	н	Î,	н	н
XA1263	СН3-	ci_ci	Н	Ŷ,	Н	Н
XA1264	снз-	a-{∑}-⊰	Н	l,	н	Н
XA1265	CH3-	ia ·	н	Ĵ,	н	Н
XA1266	СН3	a ⊜_a	н	l,	н	Н
XA1267	СН3-	a a—{}→	H	l,	н	н
XA1268	СНЗ-	a a	н	l,	н	н
XA1269	СНЗ-	H <sub>3</sub> CO_OCH <sub>3</sub>	Н	٩,	н	Н
XA1270	СН3-	OCH <sub>3</sub> H <sub>3</sub> CO-⟨□⟩→	Н	<u>گ</u>	н	н
XA1271	снз-	OCH <sub>3</sub> → H <sub>3</sub> CO	н	Ŷ,	Н	н





No.	R1	R2	R3	R4	R5	R6
XA1272	снз-	OCH <sub>3</sub> ○H OCH <sub>3</sub>	Н	Ļ,	н	н
XA1273		H₃CO H₃CO-{_}}-}	Н	l,	Н	н
XA1274	снз	H₃CQ ————————————————————————————————————	н	l,	н	Н
XA1275	СН3-	F_OCH <sub>3</sub>	Н	l,	н	Н
XA1276	СН3-	OCH <sub>3</sub>	Н	l,	н	Н
XA1277	снз-	F—COCH <sub>3</sub>	Н	Î,	Н	н
XA1278	снз-	OCH <sub>3</sub>	н	L,	н	н

_	_		
	)	Y	
١.		*	







No.	RI	R2	R3	R4	R5	R6
XA1294	СН3-	H <sub>3</sub> CO-{\bigcirc}	Н	<b>L</b> ,	Н	Н
XA1295	СН3	H <sub>3</sub> CO	н	<u>گ</u>	н	Н
XA1296	СН3-	CI, H₃CO-{\}{	н	Å,	н	н
XA1297	СН3-	F_CH <sub>3</sub>	Н	Ŷ,	н	н
XA1298	СН3-	CH₃ F—{}	Н	Å,	н	Н
XA1299	снз–	CH <sub>3</sub> F	н	Î,	н	Н
XA1300	снз-	CH <sub>3</sub>	н	Ŷ,	Н	Н

(	

No.	RI	R2	R3	R4	R5	R6
XA1301	СН3-	H <sub>3</sub> C F—{}	н	Ŷ,	н	н
XA1302	снз-	H₃C F	н	Ŷ,	Н	н
XA1303	снз-	H₃C_F	Н	l,	н	н
XA1304	CH3-	H₃C-⟨=}	Н	Ŷ,	Н	н
XA1305	СН3-	H₃C F	Н	l,	Н	н
XA1306	снз-	H <sub>3</sub> C	Н	2,	Н	н
XA1307	снз-	Br OCH <sub>3</sub>	н	Î,	Н	н
XA1308	снз-	Br—{OCH <sub>3</sub>	Н	Î,	н	н
XA1309	снз-	OCH <sub>3</sub>	н	l,	н	н
XA1310	снз-	OCH <sub>3</sub> Br	Н	l,	н	н
XA1311	СНЗ-	H₃CO Br—⟨¯¯)—}	Н	l,	Н	н
XA1312	снз-	H <sub>3</sub> CO Br	Н	Ŷ,	Н	
XA1313	СН3-	H₃CO_Br	н	l,	н	Н
XA1314	снз-	H₃CO-⟨¯¯);	Н	l,	Н	н
XA1315	снз-	H₃CO Br	н	l,	Н	Н





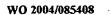
No.	R1	R2	R3	R4	R5	R6
XA1316		Br, H₃CO-⟨}—;	Н	<u></u> ,	Н	Н
XA1317	снз-	H <sub>3</sub> CO >	Н	l,	Н	н
XA1318	снз-	OCH <sub>3</sub>	Н	l,	Н	н
XA1319	снз-	CN-C-OCH3	H	Î,	Н	н
XA1320	снз-	H <sub>3</sub> CO N	Н	Ŷ,	н	н
XA1321	снз-	H³CO H³CO	H	l,	н	Н
XA1322	снз-	CN OCH3	н	١,	н	Н

						•
No.	RI	R2	R3	R4	R5	R6
XA1323	СН3-	F-(2)-1 F-	Н	l,	Н	Н
XA1324	снз-	OCH <sub>3</sub>	н	Å,	Н	Н
XA1325	снз-	H₃CO-{\}_{F}	н	Ŷ,	Н	н
XA1326	СН3-	OCH <sub>3</sub> F-{_}} OCH <sub>3</sub>	Н	Å,	н	Н
XA1327	CH3-	OCH <sub>3</sub> H <sub>3</sub> CO-(_)} OCH <sub>3</sub>	Н	Î,	Н	Н
XA1328	СН3-	a c⊢	Н	Î,	н	Н
XA1329	снз-	OCH <sub>3</sub> ci—{_}} ci	Н	Ŷ,	Н	Н
XA1330	снз-	;co-{_}; Ci	Н	l,	Н	Н
XA1331	СН3-	OCH <sub>3</sub>	н	l,	н	Н
XA1332	СН3-	H <sub>3</sub> CO-{_}-} OCH <sub>3</sub>	Н	Ŷ,	Н	н
XA1333	СН3-	OCH <sub>3</sub>	H	Ŷ,	Н	н
XA1334	CH3-	H³CO	н	Ŷ,	Н	Н
XA1335	CH3-	H³CO-{_}-{_}-1	н	Ŷ,	Н	н
XA1336	снз-	OCH <sub>3</sub> \t	н	l,	Н	Н
XA1337	СН3	H <sub>3</sub> CQ ,	Н	Ŷ,	Н	Н





No.	R1	R2	R3	R4	R5	R6
XA1338	снз-	н₃со-{∑-{С}`	н	Ŷ,	н .	Н
XA1339	СН3	OCH <sub>3</sub>	Н	Î,	н	н
XA1340	СН3-	H <sub>3</sub> CO	Н	L,	н	н
XA1341	СН3-	н₃со-⟨_>-{	Н	L,	н	Н
XA1342	CH3-	F C	Н	Î,	н	Н
XA1343	СН3-	F	н	Ŷ,	н	Н
XA1344	снз-		н	l,	Н	Н



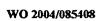


			·	<u>.                                    </u>		
No.	R1	R2	R3	R4	R5	R6
XA1345	снз-	Ø-Ø	н	Ŷ,	н	Н
XA1346	снз-	<b>\(\bar{C}\)</b>	н	Ŷ,	Н	н
XA1347	снз-	F-()-()	н	<u>گ</u>	н	Н
XA1348	сн3-	QÓ	н	بُر	н	н
XA1349	СН3-	<b>₽</b> ₽	н	بُ	н	н
XA1350	снз-	FO-Q	н	گ,	н	н
XA1351	снз-	CQ	н	Ĵ,	н	н
XA1352	снз	CCC'	Н	بُر	Н	н
XA1353	СН3-	CT;	н	Å,	н .	н
XA1354	СН3-		Н	<u></u>	Н	H
XA1355	снз-	Č <sub>r</sub>	н	گہ	Н	н
XA1356	снз-	TON .	н	بُر	н	н
XA1357	снз	,CI)	Н	Ů,	н	н
XA1358	снз–	Ţì	н.	Ŷ,	н	н
XA1359	снз-		н	١,	н	н
XA1360	снз–		Н	<u>گ</u>	н	н
XA1361	снз–	j.	н	Î,	н	н
XA1362	снз-	TOO .	н	<u></u>	н	н
XA1363	снз–	,CCC)	н	l,	н	н
XA1364	снз-	Ť.	н	گہ	н	н
XA1365	снз-	CZ;-	Н	l,	н	н
XA1366	СН3-	O.	н	١,	н	Н





No.	Ri	R2	R3	R4 -	R5	R6
		7				
XA1367	СН3-		Н	Ŷ,	Н	Н
XA1368	снз-	TOP	н	<b>A</b> ,	Н	н .
XA1369	CH3-	,CCQ	н	Ŷ,	н	н
XA1370	снз-	Ţ?	н	Ŷ,	н	н
XA1371	CH3-		Н	Ŷ,	н	н
XA1372	CH3	- Tr	Н	Ŷ,	Н	н
XA1373	снз–		н	Ŷ,	Н	н
XA1374	снз–	,Cir	н	Ĵ,	н	н
XA1375	снз-	Ţ	н	Ŷ,	Н	н
XA1376	СН3-		н	٤	Н	н
XA1377	снз-	Č,	Н	Å,	н	Н
XA1378	снз-	TON .	Н	Ĵ,	н	Н.
XA1379	СН3-	(I)	н	Ŷ,	н	н
XA1380	снз–	Č,	н	Ŷ,	Н	н
XA1381	снз-	TO!	н	Ŷ,	Н	Н
XA1382	снз	, CC	н	Ŷ,	Н	н
XA1383	снз-	Ţ\$	н	. Å,	н	Н
XA1384	снз-	O.S.	н	ŝ,	Н	Н
XA1385	СН3-	Ţ,	н	Ŷ,	Н	н
XA1386	снз-	'O's	н	ŝ,	Н	Н
XA1387	снз-	, CI\$	н	Î,	Н	Н
XA1388	снз-	S S	Н	<u>گ</u>	н	н





No.	Ri_	R2				
140.	<del> </del>	72	R3	R4	R5	R6
XA1389	CH3-	CT'r	н	<u>}</u> ,	н	н
XA1390	снз-	Č.	н	بُر	н	н
XA1391	снз-	<b>TO3</b> 1	н	Î,	н	н
XA1392	снз-	,CT	н	ئے	н	н
XA1393	СН3-	Ĩ3	н	Î,	Н	н
XA1394	снз–	Q's	н	l,	н	н
XA1395	снз-	Ţ,	н	Î,	н	н
XA1396	снз-	"CI"	н	2,	Н	н
XA1397	снз~	, CT3N	Н	3	н	н
XA1398	снз-	Ĩ,	н	Î,	н	н
XA1399	СН3-	Ţ,	н	2,	н	н
XA1400	снз-	,CCC	Н	Î,	н	Н
XA1401	СН3-	(C)	н	Î,	Н	н
XA1402	СН3-	Ō;	Н	گ,	Н	Н
XA1403	снз-	снз-	СН3-	н	Н	н
XA1404	снз-	СНЗСН2-	CH3-	н	н	Н
XA1405	СН3-	<b>∼</b> \	снз-	н	н	н
XA1406	СН3-	Y	снз-	н .	н	н
XA1407	СН3-	<b>\\\\</b>	снз-	н	н	н
XA1408	снз-	人、	снэ-	н	н	н
XA1409	СН3-	$\gamma$	снз-	н	н	Н
XA1410	СН3-	丫	СН3-	н	н	н





No.	R1	R2	R3	R4	R5	Tea
XA1411		~~~	снз-	Н	Н	R6 H
XA1412	СН3-	Y~~	СН3-	н	н	н
XA1413	снз-	X.	CH3-	Н	н	н
XA1414	снз-	7	CH3-	Н	Н	н
XA1415	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	CH3-	н ,	Н	н
XA1416	СН3-		снз-	н	н	н
XA1417	СН3-	~~~	снз-	н	Н	н
XA1418	снз–	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	снз-	Н	н	н
XA1419	снэ–	n-C8H17	СН3-	н	н	н
XA1420	СН3	L	СН3-	н	н	Н
XA1421	СН3	Q	снз-	н	н	н
XA1422	СН3-		СН3-	н	н	н
XA1423	СН3-		СН3-	н	н	н
XA1424	снз-	$\triangleright$	СН3-	н	Н	н
XA1425	СН3-	$\Diamond$	СН3-	н	н	н
XA1426	СН3		СН3-	н	н	н
XA1427	СН3-	$\bigcirc$ -1	СН3-	Н	н .	н
XA1428	снз-		СН3-	н	н	н
XA1429	СН3-	<b>○</b> -1	CH3-	н	н	н
XA1430	снз–		CH3-	н	Н	н
XA1431	СН3-		СН3-	н	н	н
XA1432	СН3-	<b>₽</b>	CH3-	Н	н	Н



No.	R1	R2	R3	R4	R5	lee
		K	-	1,14	- FG	R6
XA143:	3 CH3-		снз-	н	н	н
XA1434	СН3-		CH3-	н	н	н
XA1435	CH3-	F-()>-1	СН3-	н	н	Н
XA1436	СН3	F-On-	CH3-	н	н	н
XA1437	CH3-	CI	СН3-	н	н	Н
XA1438	снз-	CI	СН3-	н	н	н
XA1439	CH3-	c <del></del>	СН3-	н	н	н
XA1440	снз-	C⊢ <b>(</b> )→{	СН3-	Н	н	н
XA1441	снз-	CI()···{	СН3-	н	н	Н
XA1442	снз-	Br	СН3-	н	н	Н
XA1443	СН3-	Br.	СН3-	Н	н	н
XA1444	снз-	Br-{\rightarrow}-{	СН3-	н	н	н
XA1445	снз-	Br-C>-{	CH3-	н	н	н
XA1446	снз-	Br{\_\mu_\mi_\mi_	снз-	н	н	н
XA1447	снз-		CH3-	н	н	н
XA1448	снз-		СН3-	н	н	н
XA1449	снз-		СН3-	Н	Н	н
XA1450	снз-	CH <sub>3</sub>	СН3	Н	н	Н
XA1451	CH3-	H₃C 	снз-	н	Н	н
XA1452	снз-	H³C-{_}	снз-	н	н	Н
XA1453	снз–	C <sub>2</sub> H <sub>5</sub> -{	снз-	Н	н	н
XA1454	снз-	'n-C³H <sup>λ</sup> -{}-{	СН3-	н	н	н



No.	R1	R2	R3	R4	R5	R6
XA1455	СН3-	n-C <sub>4</sub> H <sub>9</sub> -{_}-{	СН3-	н	н	н
XA1456	снз-	OH	СН3-	Н	н	Н
XA1457	снз-	HO	СН3-	н	Н	н
XA1458	снз-	HO-{}-{	снз-	н .	н	н
XA1459	снз-	OCH₃	снз-	н	н	Н
XA1460	снз-	H₃CO <b>(_)</b> —∤	снз-	н	н	н
XA1461	СН3-	H <sub>3</sub> CO-{}-{	СН3-	н	Н	н
XA1462	снз-	H3CO-{}-{	снз-	н	н	н
XA1463	снз-	H3CO-{_}\{	снз-	н	Н	Н
XA1464	снз-	OC <sub>2</sub> H <sub>5</sub>	СН3-	н	Н	н
XA1465	снз-		СН3-	н	н	н
XA1466	снз-		снз–	н	н	н
XA1467	СН3-	n-C <sub>3</sub> H <sub>7</sub> O-{}-{	снз-	Н	н	н
XA1468	СН3-		снз-	н	н	н
XA1469	СН3-	NO <sub>2</sub>	СН3	н	н	н
XA1470	СН3-	O <sub>2</sub> N	снз-	н	н	н
XA1471	СН3-	O <sub>2</sub> N-{_}	СН3-	н	Н	н
XA1472	СН3-	CN	CH3-	н .	н	Н
XA1473	снз-	NC.	снз-	Н	н	н
XA1474	снз-	NC-{}-{	снз-	н	н	н
XA1475	снз-	NH <sub>2</sub>	CH3	н	н	н
XA1476	снз-	H <sub>2</sub> N	СН3-	н	н	н

No.	R1	R2.	R3	R4	R5	R6
XA1477	снз-	H <sub>2</sub> N-⟨\bar{\bar{\bar{\bar{\bar{\bar{\bar{				Н
XA1478	СН3-		СН3-	н .	н	н
XA1479	СН3-	Me <sub>2</sub> N	СН3-	Н	н	н
XA1480	CH3-	Me₂N-{}	СН3-	н	Н	н
XA1481	снз-		снз-	н	н	н
XA1482	СН3-		снз-	н	Н	н
XA1483	снз-		СН3-	н	н	н
XA1484	снз-		СН3-	н	н	н
XA1485	снз–		снз-	н	Н	н
XA1486	СН3-		снз-	н	н .	н
XA1487	СН3-		СН3-	н	н	н
XA1488	снз-		СН3-	н	н	н
XA1489	CH3-	<b>€_h-{</b> }-1	СН3-	н	н	н
XA1490	CH3-	H³CN N-{	снз-	н	H.	н
XA1491	снз-	H³CN N-{}	снз-	н	н	н
XA1492	снз-	H3CN_N-{_}}-1	СН3-	н .	н	Н
XA1493	СН3-	OCH₃ F—{}	СН3-	Н	н	н
XA1494	снз-	OCH <sub>3</sub>	СН3-	н	н	н
XA1495	снз-	OCH <sub>3</sub>	СН3-	н	н	н
XA1496	СН3-		снз-	н	н	н
XA1497	сн3-	OD,	снз-	Н	н	н
XA1498	снз-	снз-	н	н	снз-	н



No.	R1	R2	R3	R4	R5	R6
XA1499	снз-	СН3СН2-	Н	Н	снз-	н
XA1500	CH3-	<b>^</b> `\	н ·	н	снз-	н
XA1501	снз-	Y	н	н	снз-	н
XA1502	снз–	<b>~</b> ~\	Н	н	снз-	н
XA1503	CH3-	人、	н	н	сн3-	H
XA1504	CH3-	$\gamma$	Н	н	снз-	H
XA1505	СН3-	丫	Н	н	снз-	Н
XA1506	снз-	^^\	Н	н	СН3-	н
XA1507	снз-	$\rightarrow$	Н	н	снз-	н
XA1508	СН3-	\\\\	н	н	СН3-	н
XA1509	снз–	分	н	н	сна-	н
XA1510	снз–	<b>&gt;&gt;&gt;&gt;</b>	н	н	снз-	н
XA1511	снз–		н	н	снз-	н
XA1512	снз–	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н	снз-	н
XA1513	снз–	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н	снз-	н
XA1514	СН3	л-С8Н17	H	н	С <b>Н3</b> -	н
XA1515	снз–	L~~~	н	н	снз-	Н
XA1516	СН3	Q	н	н _	СН3-	н
XA1517	снз-		н	н	снз-	Н
XA1518	СН3-		Н	Н	снз-	н
XA1519	снз–	<b>⊳</b> ⊣	н	н.	снз-	н
XA1520	СН3-	·	н	н	снз-	н



No.	R1	R2	R3	R4	R5	R6
XA1521	снз-	$\bigcirc$	Н	н	СН3-	н
XA1522	снз-	$\bigcirc$ $\dashv$	Н	н	снз-	н
XA1523	снз	$\bigcirc \dashv$	Н	н	СН3-	н
XA1524	снз-		Н	н	СН3-	н
XA1525	снз-		Н	н	снз-	н
XA1526	снз-	<b>_</b> m4	Н	Н	снз-	н
XA1527	снз-		Н	н	СН3-	н
XA1528	снз–	F	Н	н	снз-	н
XA1529	снз-	F-()-1	Н	н	снз-	н
XA1530	снз-	F-(-)	Н	н	снз-	н
XA1531	снз-	F———III-{	Н	н	СН3-	н
XA1532	снз-	CI C)	н	н	снз-	н
XA1533	СН3-	CI \	н	н	снз-	н
XA1534	СН3-	c⊢ <b>∕</b>	Н	Н	снз-	н
XA1535	сн <del>з-</del>	c⊢ <b>(</b> )—(	н	Н	снз-	н
XA1536	снз-	CI—(	н	Н	снз-	Н
XA1537		Br ⊘–∤	н	н	снз-	н
XA1538	снз-	Br.	н	н	снэ-	н
XA1539	снэ–	Br- <b>(</b> ){	Н	н	снз-	н
XA1540	снз-	Br- <b>(</b> ){	н	н	снз-	Н
XA1541	снз-	Br{\_\mid_\mid	н	н .	снз-	н
XA1542	снз-		Н	н	снз-	Н

XA1563

XA1564

снз~

СН3-

NO<sub>2</sub>

No.	R1	R2	R3	R4	R5	R6
XA1543	СН3-		н	Н	CH3-	н
XA1544	снз-		н	н	CH3-	н
XA1545	снз-	CH₃	н	н	CH3-	н
XA1546	снз–	H <sub>3</sub> C	н	н	снз-	н
XA1547	снз	H3C-{	н	н	CH3-	н
XA1548	снз-	C <sub>2</sub> H <sub>5</sub> -{_}-{	н	Н	CH3-	Н
XA1549	снз-	л-С <sub>3</sub> Н <sub>7</sub> ⟨}-{	н	н	снз-	н
XA1550	снз-	n-C <sub>4</sub> H <sub>9</sub> -	н	н	снз-	н
XA1551	снз-	OH OH	н	Н	снз-	н
XA1552	CH3-	HO	н	Н	снз-	н
XA1553	CH3-	HO-{}-{	н	н	снэ–	н
XA1554	CH3-	OCH <sub>3</sub>	н	н	снз-	н
XA1555	снз-	H <sub>3</sub> CO	н .	н	CH3-	н
XA1556	снз-	H₃CO-{}	н	Н	CH3-	н
XA1557	снз-	H3CO-{}-{	н	Н	снэ-	н
XA1558	снз-	H³CO-{\rightarrow}\!\!	н	н	CH3-	н
XA1559	снз-	OC <sub>2</sub> H <sub>5</sub>	н	н	снз-	н
XA1560	СН3-	C₂H₅O —}-{	н	н	CH3-	н
XA1561	СН3-	C <sub>2</sub> H <sub>5</sub> O-	н	н	CH3-	н
XA1562	CH3-	n-C <sub>3</sub> H <sub>7</sub> O-{}-{	н	н	CH3-	н

н

н

н

снз-

СН3-

Н

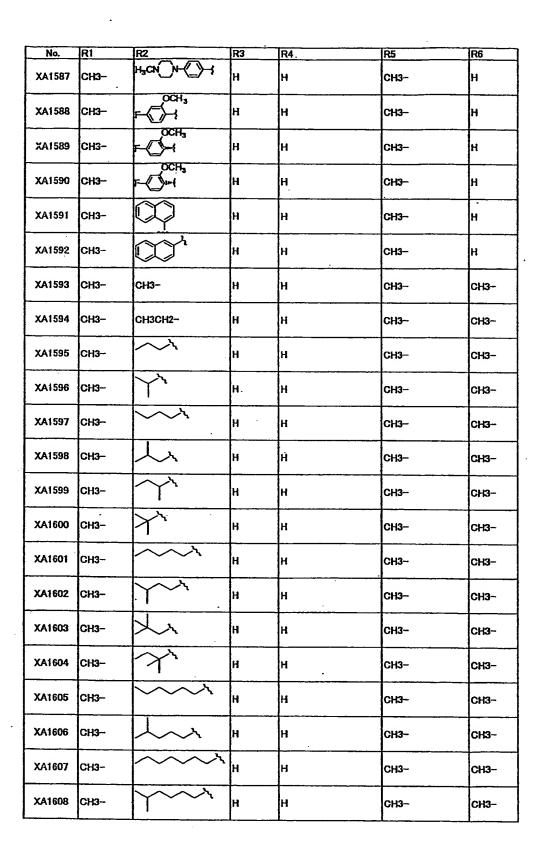
н



No.	R1	R2	R3	R4	R5	R6
XA1565	снз-	O <sub>2</sub> N	н	н	снз–	н
XA1566	СН3-	02N-{}-{	н	н	СН3-	н
XA1567	СН3-	CN CN	н	н	снз-	Н
XA1568	СН3-	NC	н	Н	СН3	н
XA1569	снз-	NC-()-1	н	н	снз-	Н
XA1570	снз-	NH <sub>2</sub>	н	н	СН3-	н
XA1571	СН3-	H <sub>2</sub> N	н	H	СН3-	н
XA1572	СН3-	H <sub>2</sub> N-(T)	н	Н	снз-	н
XA1573	снз-	NMe <sub>2</sub>	Н	Н	СН3-	Н
XA1574	снз-	Me <sub>2</sub> N ☐  —  —  —  —  —  —  —  —  —  —  —  —	Н	н	снз-	н
XA1575	снз-	Me <sub>2</sub> N-	н	н	СН3~	н
XA1576	СН3-		н	н	CH3-	н
XA1577	СН3-		н	Н	СН3-	Н
XA1578	снз-	CH-C>-1	Н	н	СН3-	н
XA1579	снз–		Н	н	CH3-	н
XA1580	снз-		н	н	CH3-	Н
XA1581	снз-		н	н	снз-	н
XA1582	снз-		Н	н .	СН3-	н
XA1583	снз-	O+Q	Н	Н	CH3-	н
XA1584	СН3-		н	н	снз-	н
XA1585	СН3-	H³CN_N-	н	н	СН3-	н
XA1586	СН3	H³CN_N-{\bigs_}	н	н	СН3-	н

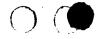








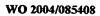
No.	R1	R2	R3	R4	R5	R6
XA1609	снз-	n-C8H17-	н	н	СН3-	СН3-
XA1610	СН3-	J.,,,	н	Н	СН3-	СН3
XA1611	сна-	Q	Н	Н	снз-	СН3-
XA1612	снз-		н	Н	снз-	СН3-
XA1613	снз-		н	н	снз-	CH3-
XA1614	СН3-	D{	н.	н	снз-	CH3
XA1615	снз	$\Diamond$ -1	H <sub>.</sub>	Н	снз-	СН3-
XA1616	снз-	$\bigcirc$	Н	Н	снэ-	CH3-
XA1617	снз–	$\bigcirc$ $\dashv$	Н	Н	снз–	CH3-
XA1618	СН3-	$\bigcirc$ $\dashv$	н	н	снз-	СН3-
XA1619	СН3-		н.	Н	СН3-	снз-
XA1620	СН3-	$\bigcirc$ $\vdash$	Н	Н	СН3-	CH3-
XA1621	CH3-	<b></b>	н	н	СН3-	снз-
XA1622	СН3		Н	Н	СН3-	СН3-
XA1623	снз-		Н	н	СН3-	СН3
XA1624	СН3	F	Н	н	снз-	CH3-
XA1625	снз-	F-()	Н	н	СН3-	снз–
XA1626	снз-	F-{\rightarrow}n.{	н	н	СН3-	СН3
XA1627	СН3-	CI C	н	н	СН3-	CH3-
XA1628	СН3-	Ci	Н	Н	снз–	снз-
XA1629	СН3-	c <del>(-)-</del> -{	н	н	СН3-	снз–
XA1630	снз-	c⊢ <b>∕</b>	H	Н	СН3-	СН3



No.	R1	Ing.	lee .	- Im-		- r
NO.	KI	R2	R3	R4	R5	R6
XA1631	снз-	CI-{_}\{	н	н	снз-	снз-
XA1632	снз-	Br	н	н	снз-	СН3-
XA1633	снз-	Br.	н	н	снз-	снз-
XA1634	снз-	Br-C}-{	н	н	снз-	снз-
XA1635	снз-	Br—()—i	Н	Н	снз-	снз-
XA1636	снз-	Br————	н .	Н	снз-	снз-
XA1637	снз-		н	н	снз-	снз-
XA1638	снз-		Н	н	снз-	снз-
XA1639	снз-		н	Н	СН3-	CH3-
XA1640	снз-	CH <sub>3</sub>	н	H	снз-	снз-
XA1641	снз-	H <sub>3</sub> C	H	Н	снз-	снз-
XA1642	СН3-	H³C-{}	н	н	снз-	снз–
XA1643	снз–	C₂H₅-⟨}-	Н	н	снз-	снз–
XA1644	снз-	n-C₃H <sub>7</sub> -{	н	н	СН3-	СН3-
XA1645	снз-	n-C₄H <sub>9</sub> -∕	н	Н	снз-	СН3-
XA1646	снэ-	OH OH	н	н	СН3-	снз-
XA1647	снз-	HO —	н	н	CH3-	CH3-
XA1648	снз-	HO-{_}-	н,	н	СН3-	снз-
XA1649	снз-	OCH₃	н	н	снз-	снэ-
XA1650	снз-	H₃CO ————————————————————————————————————	н	н	снз-	снз-
XA1651	снз-	H₃CO- <b>{</b> }-{	н	н	снз-	снз-
XA1652	снз-	H₃CO- <b>(</b> )~{	н	Н	снз-	снз-



No.	R1	R2	R3	R4	R5	Inc
		H <sub>3</sub> CO-( )n-(	1.0	IV4	NO .	R6
XA1653	CH3-		Н	Н	CH3-	снз-
XA1654	снз-	OC <sub>2</sub> H <sub>5</sub>	н	н	снз-	снз-
XA1655	СНЗ	C₂H₅O	н	н	СН3-	снз-
XA1656	CH3-		Н	н	снз-	снз-
XA1657	Сн3-	n-C <sub>3</sub> H <sub>7</sub> O- <b>⟨</b> _}-{	Н	Н	снз-	
XA1658	СН3-	n-C₄H <sub>9</sub> O-⟨}	Н	н	СН3-	снз-
XA1659	СН3-	NO <sub>2</sub>	Н	н	снз-	снз–
XA1660	снз-	O <sub>2</sub> N	н	н	СН3-	снз-
XA1661	снз-	021-	н	н	СН3-	снз-
XA1662	снз-	CN	Н	н	СН3-	снз-
XA1663	СН3-	NC .	н	н	СН3-	снз-
XA1664	CH3-	NC-{}-{	н	н	снз-	снз–
XA1665	снз-	NH <sub>2</sub>	н	н	CH3-	снз-
XA1666	СН3-	H <sub>2</sub> N	Н	н	снз-	снз
XA1667	СН3-	H <sub>2</sub> N-{}-{	Н	Н	снз-	снз-
XA1668	снз-		Н	н	снз-	снз-
XA1669	СН3-	Me <sub>2</sub> N	Н	Н	снз-	снз-
XA1670	снз-	Me <sub>2</sub> N-{	н	Н	снз-	снз-
XA1671	снз–		Н	н	СН3-	снз-
XA1672	СН3-		н	Н	СН3-	снз-
XA1673	СН3		н	н	CH3-	СН3-
XA1674	снз–	O+(2)	н	Н	CH3-	CH3-



1		
(	)	
٦.	1	

<del></del>						
No.	R1	R2	R3	R4	R5	R6
XA1675	СН3-	(n<)	н	н	снз-	СН3-
XA1676	СН3-	(N-()-1	н .	н	снз-	СН3-
XA1677	СН3-	Or	н	н	СН3-	СН3-
XA1678	СН3-	<b>○~</b> ○	н	н	снз-	СН3-
XA1679	снз-	<b>○ખ</b> ○→	н	н	снз-	CH3-
XA1680	снз-	H3CN N-	н	н	снэ-	CH3-
XA1681	снз-	H3CN N-C	н	н	СН3-	CH3-
XA1682	снз-	H²CN_N-{}-{	н	н	снз-	СН3-
XA1683	снз-	OCH₃ F-{}	н	н	снз-	СН3-
XA1684	снз-	OCH <sub>3</sub>	Н	н	СН3	СН3-
XA1685	снз-	OCH <sub>3</sub>	н	Н	снз-	СН3-
XA1686	СН3-		н	Н	СН3-	СН3-
XA1687	снз-	CC,	Н	Н	СН3-	СН3-
XA1688	снз-	снз-	н	СН3-	снз-	СН3-
XA1689	снз-	СНЗСН2-	н	СН3-	снз-	снз-
XA1690	CH3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	снз-	снз-	снз-
XA1691	снз-	<u> </u>	н	снз-	CH3-	СН3-
XA1692	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	СН3-	СН3-	СН3-
XA1693	снз-	人、	н	снз-	снз-	СН3-
XA1694	снз-	~	н	снз-	СН3-	снз-
XA1695	СН3-	_'	н	СН3-	СН3-	CH3
XA1696	СН3-	^~\`\	Н	снз-	СН3-	CH3





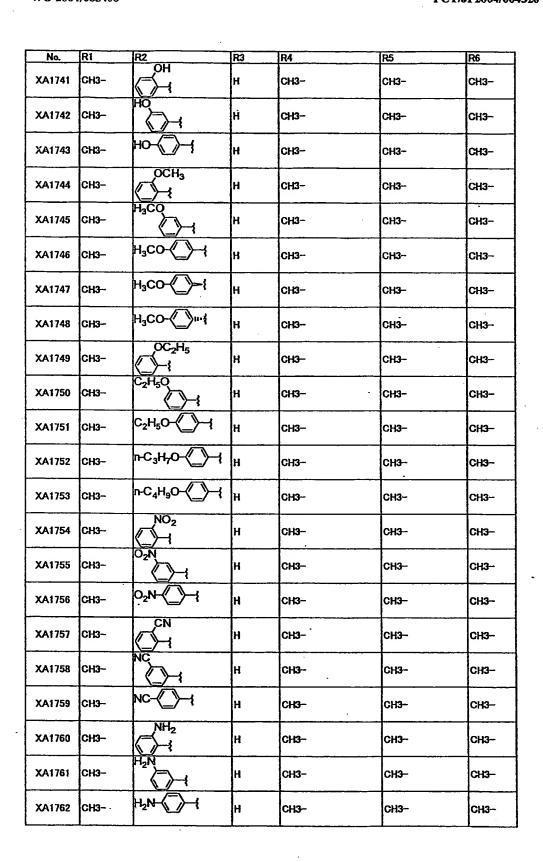
No.	R1	R2	R3	R4	R5	R6
XA1697	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	снз–	СН3	снз-
XA1698	снз-	×x	н	снз-	СН3-	СН3-
XA1699	снз-	7	н	снз-	СН3-	СН3-
XA1700	снз-	<b>~~~</b>	Н	снз–	снз-	снз-
XA1701	снз-	<u></u>	Н	СН3-	СН3-	С <b>Н3</b> –
XA1702	снз-	~~~``	н	СН3-	СН3	снз-
XA1703	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н .	СН3-	СН3-	снз-
XA1704	снз–	n-C8H17-	н	СН3-	снз–	снз-
XA1705	снз-	人~~	н	СН3-	СН3-	СН3-
XA1706	СН3-		Н	СН3-	СН3	снз-
XA1707	снз-		Н	СН3	снз-	снз-
XA1708	снз-		H	CH3-	снз-	СН3-
XA1709	снз-	<b>▷</b> →	Н	CH3-	СН3-	снз-
XA1710	снз-	$\Diamond$ -I	H	CH3	снз-	снз-
XA1711	СН3-	$\bigcirc$	н	СН3	снз–	СН3-
XA1712	снз–	. 🖂	н	CH3-	СН3-	CH3-
XA1713	снз	H	H	CH3	снз-	СН3-
XA1714	СН3-		н	СН3-	СН3-	СН3
XA1715	снз-		Н	CH3-	СН3-	СН3-
XA1716	снз-	<b></b>	Н	СН3-	снз-	СН3
XA1717	СН3-	F	H	СН3	снз-	CH3-
XA1718	CH3-		Н	снз-	СН3-	снз-



1 No	R1	Inc		<del></del>		
No.	IKI .	R2	R3	R4	R5	R6
XA1719	СН3-	F-(_)-1	н	снз–	снз-	CH3-
XA1720	снз-		н	снз-	СН3-	СН3-
XA1721	снз–	F-()(	н	СН3	СН3-	снз-
XA1722	снз-	CI	н	снз	СН3-	снз-
XA1723	снз-	CI	Н	снз–	СН3-	CH3
XA1724	снз-	CH	н	CH3	СН3-	СН3-
XA1725	снз–	CH	н	СН3	снз-	снз-
XA1726	снз-	CI-{\bigs_m\	Н	снэ-	СН3-	снз-
XA1727	снз–	Br	Н	снз-	CH3-	CH3-
XA1728	снз–	Br.	н	СН3-	СН3-	CH3-
XA1729	снз-	Br-{}-{	н	снз-	СН3-	CH3-
XA1730	снз-	Br—{}	Н	снз-	СН3	СН3-
XA1731	снз	Br————	н	снз-	СН3-	СН3
XA1732	снз–	<b>◯</b> -₁	н	снз-	СН3-	СН3-
XA1733	снз-	<u></u>	н	СН3-	СН3-	СН3-
XA1734	снз-	<b>⊢</b>	н	снз-	СН3-	СН3-
XA1735		CH₃ ◯≻⊰	н	СН3-	снз-	СН3-
XA1736	СН3	H₃C	Н	CH3-	СН3-	СН3-
XA1737	СН3-	H³C- <b>{</b> }{	н	снз-	снз-	СН3-
XA1738	СН3-	C <sub>2</sub> H <sub>5</sub> -{}-{	H	снз-	СН3-	СН3-
XA1739	СН3-	n-C₃H <sub>7</sub> -{_}{	н	СН3-	СН3-	CH3-
XA1740	СН3-	ŋ-C₄H <sub>9</sub> -⟨}-	Н	СН3-	СН3-	СН3-









1		

No.	RI	IDO	Too		les.	<del></del>
NO.	KI	R2 NMe <sub>2</sub>	R3	R4	R5	R6
XA1763	CH3-	<b>⟨</b>	Н	снз-	СН3-	CH3-
XA1764	СН3-	Me <sub>2</sub> N	Н	СН3-	СН3-	снз-
XA1765	снз-	Me <sub>2</sub> N-(	н	снэ	СН3-	снз-
XA1766	снз-		н	снз-	снз-	СН3-
XA1767	снз-		н	снз-	СН3-	снз-
XA1768	СН3-		н	снз-	СН3-	снз-
XA1769	снз-		Н	СН3-	снз-	снз-
XA1770	СН3-		Н	СН3-	СН3-	СН3-
XA1771	снз-		н	СН3-	СН3-	снз-
XA1772	снз-	√	Н	СН3-	снз-	снз-
XA1773	СН3-		Н	снз-	снз-	снз-
XA1774	снз-		Н	снз-	CH3-	снэ-
XA1775	снз-	H³CN N-	Н	снз-	СН3-	сн3-
XA1776	снз-	H3CN N-	Н	СН3-	CH3-	снз-
XA1777	снз-	H3CN_N-{}-	Н	снз-	CH3-	СН3-
XA1778	СН3	F-€]}-¦	Н	СН3-	снз-	снз-
XA1779	снз–	CH3 F—⊘H	н	СН3	снз-	СН3-
XA1780	снз-	OCH <sub>3</sub>	н	СН3-	СН3-	снз-
XA1781	СН3-		н	СН3-	снз-	снз-
XA1782	снз-	CC,	н	снз–	снз-	снз-
XA1783	СНЗСН2-	СН3-	н	н	Н	н
XA1784	СНЗСН2-	СН3СН2-	н	Н	н	н







No.	R1	R2	R3	R4	R5	R6
XA1785	Снзсн2-	~^\	н	н	н	Н
XA1786	снзсн2-	Y	Н	н	н	н
XA1787	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н	н	Н
XA1788	снзсн2-	人工	н	Н	Н	Н
XA1789	снзсн2-	~\^`	Н	н	н	н
XA1790	СНЗСН2-	·	Н	н	н	н
XA1791	СН3СН2-	~~\\	н	н	н	н
XA1792	СНЗСН2-	<b>\</b>	н	н	н	Н
XA1793	снзсн2-	<u> </u>	н	н	н	H .
XA1794	СНЗСН2-	7	н	н	н	н
XA1795	СН3СН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	н	н	н
XA1796	СНЗСН2-		н	н	н	н -
XA1797	СН3СН2-	^^^\	н	Н	Н	н
XA1798	СН3СН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н	н	Н
XA1799	СНЗСН2-	n-C8H17-	Н	Н	н	Н
XA1800	СН3СН2-	<u> </u>	н	н	Н	н
XA1801	СНЗСН2-	Qi_	Н	н	Н	H
XA1802	СН3СН2-		н	н	Н	Н
XA1803	СН3СН2-	Q	Н	Н	Н	Н
XA1804	СН3СН2-	$\triangleright$	н	Н	н	н
XA1805	СНЗСН2-	$\Diamond$ -1	н	Н	н	н
XA1806	СНЗСН2-	$\bigcirc \dashv$	н	н	Н	н



	1=5					50
No.	R1	R2	R3	R4	R5	R6
XA1807	снзсн2-	$\bigcirc$ $\dashv$	н	H	н	Н
XA1808	снзсн2-	$\bigcirc$ $\dashv$	н	н	H	н
XA1809	снзсн2-	$\bigcirc$	н	н	Н	н
XA1810	снзсн2-		н	н	н	н
XA1811	снзсн2-	<b>⊘</b> m4	н	н	Н	Н
XA1812	снзсн2-		н	н	н	Н
XA1813	СНЗСН2-	F	Н	н	н	н
XA1814	СНЗСН2-	F-()-1	Н	Н	н	н
XA1815	СН3СН2-		H	Н	н	н
XA1816	СН3СН2-	F	н	Н	Н	н .
XA1817	СНЗСН2-	CI C)-i	н	н	н .	Н
XA1818	СНЗСН2-	CI ———	Н	Ĥ	н	н
XA1819	СН3СН2-	c⊢ <b>(</b> )–₁	Н	н	Н	н
XA1820	СНЗСН2-	c⊢ <b>(</b> )~i	н	н	н	Н
XA1821	СН3СН2-	CI()···-{	н	н	н	н.
XA1822	СНЗСН2-	Br S−{	н	н	Н	Н.
XA1823	снзсн2-	Br.	н	н	н	н
XA1824	снзсн2-	Br─∰	Н	н	н	н
XA1825	СНЗСН2-	Br—{_}_{}	н	н	н	н
XA1826	СНЗСН2-	Br-{_}m-{	н	н	н	н
XA1827	СНЗСН2-		н	Н	н	н
XA1828	снзсн2-		н	Н	Н	н

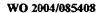




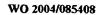
No.	R1	R2	R3	R4	R5	R6
XA1829	CH3CH2-	' \= ' '	H	Н	H	н
XA1830	СНЗСН2-	CH3	Н	.,		
AA1830		<b>⟨_</b> }⊣	n	H	H	Н
XA1831	СНЗСН2-	H₃C	н	H	н Н	н
					<u> </u>	<u> </u>
XA1832	СНЗСН2-	н₃с-{_}};	ម	H	н	н
XA1833	СНЗСН2-	C <sub>2</sub> H <sub>5</sub> ()(	н	н	н	н
		-				
XA1834	СНЗСН2-	ո-C₃H <sub>7</sub> ⟨⟩{	н	н.	н	н
XA1835	СНЗСН2-	n-C <sub>4</sub> H <sub>9</sub> <}	н	н	н	н
		ОН				
XA1836	СНЗСН2-	/= <u>C.</u>	н	н	н	н
		HO			<u> </u>	
XA1837	СНЗСН2-		н	н	н	н
		,				
XA1838	СНЗСН2-	HO-{}-}	н	н	н	н
		OCH <sub>3</sub>				
XA1839	СНЗСН2-		н	н	н	н
		H <sub>3</sub> CQ	ļ ———			
XA1840	СНЗСН2-		н	H	н	Н
		H CO-				
XA1841	СНЗСН2-	1300/	н	н	H	Н
		H.CO.				<b></b>
XA1842	снзсн2-	1,300 (, ,	H	н	Н	H
		H-CO-( )us	1			
XA1843	снзсн2-		Н	н	н	Н
		OC <sub>2</sub> H <sub>5</sub>				
XA1844	СНЗСН2-	<b>⟨</b> _ <b>⟩</b> -;	Н	н	н .	Н
VALOUE	01100110	C <sub>2</sub> H <sub>5</sub> Q				1
XA1845	СНЗСН2-	∑, ièo,	H	Н	Н	Н
XA1846	01100110				1	
AA1840	СНЗСН2-	,	н	Н	н	Н
XA1847	СНЗСН2-	n-C₃H <sub>7</sub> O-⟨}-{		.,	<b>I.</b> ,	
AA104/	UNJUNZ-		H	Н	H	Н
XA1848	СН3СН2-	n-C4H9O-{ }-{	н	н	н	н
A31040				<u></u>	<u></u>	<u></u>
XA1849	CH3CH3-	NO <sub>2</sub>		u	lu	L.
W/1043	CH3CH2-	<u> </u>	н	Н	H	Н
XA1850	CH3CH2-	O <sub>2</sub> N	Н	н	н	Н
~			<u> </u>		<u> </u>	

)	(			
,	i			

No.	RI	R2	R3	R4	R5	lpa.
<u> </u>	<del>                                      </del>	<b>6</b> 3	TG	IN4	170	R6
XA1851	СНЗСН2-	O <sub>2</sub> N-{_}}-	н	Н	н	н
XA1852	СНЗСН2-	CN CH	H	Н	н	н
XA1853	снзсн2-	NC	н	Н	н	н
XA1854	СНЗСН2-	NC-	н	н	н	н
XA1855	СНЗСН2-	NH <sub>2</sub>	н	н	H	н
XA1856	снзсн2-	H <sub>2</sub> N	н	н	н	н
XA1857	СН3СН2-	H <sub>2</sub> N-{_}}-{	н	н	н	н
XA1858	СНЗСН2-	NMe₂	н	н	Н	н
XA1859	СНЗСН2-	We <sup>5</sup> N	Н	н	н	н
XA1860	СНЗСН2-	Me₂N-{}	Н	н	Н	н
XA1861	СНЗСН2-		н	н	н	Н
XA1862	СНЗСН2-		н	н	н	н
XA1863	СНЗСН2-	(n-()-1	н -	н	н	н
XA1864	СНЗСН2-		н	н	н	н
XA1865	СНЗСН2-		н	Н	н	н
XA1866	СНЗСН2-		н	H	Н	н
XA1867	СНЗСН2-		Н	н	н	н
XA1868	СНЗСН2-		Н	н	Н	н
XA1869	снзсн2-	< <u></u> \-( <u>\</u> \-( <u>\</u> \-	Н	н	н	н
XA1870	СНЗСН2-	H³CN N-√\	Н	н .	Н	н
XA1871	СНЗСН2-	H₃CN_N-	н	н	н	н
XA1872	СНЗСН2-	H3CN/N-{}	Н	н	н	Н



No.	R1	R2	R3	R4	R5	R6
XA1873	снзсн2-	ocH, F-⟨∑}-¦	Н	н	н	н
XA1874	СНЗСН2-	осн <sub>3</sub> г-{}-{	Н	н	н	Н
XA1875	снзсн2-	OCH <sub>3</sub> F—Ou-(	Н	н	н	н
XA1876	снзсн2-		Н	Н	Н	н
XA1877	снзсн2-	CCC,	H	Н	Н	н
XA1878	СНЗСН2-	CH3-	н	снз-	н	н
XA1879	снзсн2-	СНЗСН2-	н	снз-	Н	н
XA1880	Снзсн2-	<b>∼</b> \\	н	снз-	Н	Н
XA1881	снзсн2-	$\uparrow \gamma$	н	снз-	Н	н
XA1882	снзсн2-	~~``	н	снз-	н	н
XA1883	снзсн2-	人	Н	СН3-	н	н
XA1884	снзсн2-	~~`	н	снз-	H	Н
XA1885	СНЗСН2-	Y	н	снз-	н	H ·
XA1886	СНЗСН2-	~~``	н	снз-	н	н
XA1887	снзсн2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3-	н	н .
XA1888	СНЗСН2-	\\ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	н	снз-	н	Н
XA1889	снзсн2-	7	н	СН3	Н	н
XA1890	снзсн2-	<b>\\\\\</b>	н	снз-	н	н
XA1891	СНЗСН2-	L~~	н	СН3-	н	Н
XA1892	СНЗСН2-	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3-	н	Н
XA1893	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3-	н	Н
XA1894	СНЗСН2-	n-C8H17-	н	CH3-	н	н





No.	R1	R2	R3	R4	R5	R6
140.		1	17.5	1/4	11.0	17.0
XA1895	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3	Н	н
XA1896	Снзсн2-	Qi	н	снз-	н	н
XA1897	снзсн2-		Н	СН3-	н	н
XA1898	снзсн2-	Qui	Н	снз-	Н	н
XA1899	снзсн2-	$\triangleright$	н	снз-	H	н
XA1900	СНЗСН2-	$\Diamond$	н .	снз-	н	н
XA1901	СНЗСН2-	$\bigcirc \vdash$	н	снз-	н	н
XA1902	СНЗСН2-	$\bigcirc$ $\dashv$	Н	снз-	Н	н
XA1903	СНЗСН2-	$\bigcirc$ $\vdash$	н	снз	н	н
XA1904	снзсн2-		H	СН3-	н	н
XA1905	снзсн2-		н	снз-	н	н
XA1906	снзсн2-		н	СН3-	Н	н
XA1907	СНЗСН2-		н	СН3-	Н	н
XA1908	снзсн2-	F	н	СН3	Н	н
XA1909	снзсн2-	F-()-1	н	снз-	Н	н
XA1910	СНЗСН2-	F-()	н	СН3-	н	н
XA1911	СНЗСН2-	F-()nd	н	СН3-	н	Н
XA1912	СНЗСН2-	CI C)	н	снз-	Н	Н
XA1913	СН3СН2-	CI C	н	СН3-	н	н
XA1914	снзсн2-	c⊢(_)~{	Н	СН3-	н	н
XA1915	СНЗСН2~	CI(>-	Н	СН3-	н	н
XA1916	СНЗСН2-	CH	н	снз–	н	н

No.	R1	R2	R3	R4	R5	R6
XA1917	снзсн2-	Br ∰-∤	Н	СН3-	н	н
XA1918	СНЗСН2-	Br.	Н	СН3-	н	Н
XA1919	снзсн2-	Br-{\rightarrow}-4	Н	СН3	н	н
XA1920	СНЗСН2-	Br—()	H .	CH3-	н	Н
XA1921	снзсн2-	Br—(	Н	CH3-	Н	н
XA1922	снзсн2-		H	СН3-	н	н
XA1923	СНЗСН2-		Н	СН3-	н	н
XA1924	снзсн2-		Н	СН3-	н	н
XA1925	СНЗСН2-	CH <sub>3</sub>	Н	СН3-	н	н
XA1926	СНЗСН2-	H₃C ————————————————————————————————————	Н	с <b>н</b> з	н	Н
XA1927	снзсн2-	H <sub>3</sub> C-{\rightarrow}-4	н	снз-	н	н
XA1928	снзсн2-	C <sub>2</sub> H <sub>5</sub> -{_}-{	н	снз-	н	н
XA1929	СНЗСН2-	n-C₃H <sub>7</sub> {}{	н	СН3-	н	Н
XA1930	СН3СН2-	n-C <sub>4</sub> H <sub>9</sub> -	н	СН3	н	н
XA1931	СНЗСН2-	©H ⊘H	н	сн3-	н	н
XA1932	СНЗСН2-	HO	н	СН3-	Н	Н
XA1933	снзсн2-	ю-()-	н	снз–	Н	Н
XA1934	снзсн2-	OCH <sub>3</sub>	н	снз-	н	Н
~XA1935	снзсн2-	H₃CO 	н	снз-	Н	н
XA1936	снзсн2-	H³CO- <b>⟨</b> _}-!	н	сн3-	н	н
XA1937	снзсн2-	H₃CO- <b>(</b> )-{	н	сн3-	Н	н
XA1938	снзс;н2-	H <sub>3</sub> CO-{_}m{	н	снз-	н	н



No.	R1	R2	R3	R4	R5	R6
XA1939	снзсн2-	OC <sub>2</sub> H <sub>5</sub>	н	снз-	н	н
XA1940	СНЗСН2-	C2H50	н	снз-	н	н
XA1941	СНЗСН2-		н	снз-	н	н
XA1942	СНЗСН2-	n-C <sub>3</sub> H <sub>7</sub> O-{}-{	н	CH3-	н	н
XA1943	СНЗСН2-	n-C <sub>4</sub> H <sub>9</sub> O-	Н	снз-	н	н
XA1944	СН3СН2-	NO <sub>2</sub>	н	снз-	н	н
XA1945	снзсн2-	O <sub>2</sub> N	н	снз-	н	н
XA1946	СН3СН2-	O <sub>2</sub> N-{}-{	н	снз-	н	н
XA1947	СНЗСН2-	CN ◯─┤	н .	снз-	н	н .
XA1948	СНЗСН2-	NC	н	снз	н	н
XA1949	снзсн2-	NC-{}-{	н	снз	H	н
XA1950	снзсн2-	NH <sub>2</sub>	н	СН3	н	н
XA1951	СНЗСН2-	H <sub>2</sub> N \	Н	СН3-	Н	н
XA1952	СНЗСН2-	H <sub>2</sub> N-{_}	н	СН3-	н	н
XA1953	снзсн2-	NMe₂	н	СН3-	н	н
XA1954	СНЗСН2-	Me <sub>2</sub> N	н	СН3-	Н	н
XA1955	СНЗСН2-	Me <sub>2</sub> N-{	н	СН3	н	н
XA1956	СНЗСН2-		н	СН3-	Н	н
XA1957	СНЗСН2-		н	снз-	н	Н
XA1958	снзсн2-		н	СН3-	н	н
XA1959	снзсн2-		Н	СН3	н	н
XA1960	снзсн2-		н	СН3-	н	н



No.	R1	R2	R3	R4	R5	R6
XA1961	снзсн2-		Н	снз-	н	н
XA1962	снзсн2-		н	снз-	н	н
XA1963	снзсн2-		н	снз-	н	н
XA1964	снзсн2-		Н	снз-	Н	н
. XA1965	СНЗСН2-	H3CN N-	Н	СН3-	н	н
XA1966	снзсн2-	H3CN N-C	н	СН3	н	н
XA1967	снзсн2-	H³CN_N-{\}-\	н	снз-	н	н
XA1968	снзсн2-	F-€S-4	H	снз-	Н	н
XA1969	СНЗСН2-	OCH <sub>3</sub>	н	СН3-	Н	н
XA1970	СНЗСН2-	OCH <sub>3</sub>	н	СН3-	н	н
XA1971	снзсн2-	CQ ·	н	снэ-	н	н
XA1972	снзсн2-		н	СН3-	н	н

No.	STRUCTURE
VA 4070	
XA1973	CI N N N N O CH <sub>3</sub>
XA1974	Br NN O CH <sub>3</sub>
XA1975	CH <sub>3</sub> N N N CH <sub>3</sub> O CH <sub>3</sub> O CH <sub>3</sub>
XA1976	CIH CIH N N O CH <sub>3</sub>

XA1977	CIH CIH CH CH3
XA1978	a CH <sub>N</sub> O
XA1979	
XA1980	HCI HCI HCI CH <sub>3</sub>

XA1981	HCI HCI HCI CH <sub>3</sub>
XA1982	
	Ha Ha Ha Ha CH <sub>3</sub> C N CH <sub>3</sub>
XA1983	CIH N
	CIH CIH
	CH <sub>3</sub>
XA1984	СІН
	CI N N O CH <sub>3</sub>

V44005	T
XA1985	CIH CH CH, CH, CH,
XA1986	CH CH, CH, CH, CH, CH, CH, CH, CH, CH, C
XA1987	CIH CIH CH <sub>3</sub>
XA1988	HCI HCI N CH <sub>3</sub> O



XA1989	HO HO HO P CH
XA1990	Ha Ha Ha Ha Ha Ha CH3
XA1991	Ha Ha CH
XA1992	CIH CIH CH <sub>3</sub>

XA1993	CIH CH <sub>3</sub> CH <sub>3</sub>
XA1994	CIH CIH CH <sub>3</sub> CH <sub>3</sub>
XA1995	CIH CH CH <sub>3</sub> C CH <sub>3</sub>
XA1996	CH <sub>3</sub> CiH CiH N CH <sub>3</sub> CH <sub>3</sub>





VA4007	
XA1997	CH <sub>3</sub> CH CH N N N N N N N N N N N N N N N N N N N
XA1998	CIH CIH N N N N CH <sub>3</sub>
XA1999	HCI  CIH  CIH  N  CH  CH  CH  CH  CH  CH  CH  CH  CH
XA2000	CIH CH, CH, CH,





XA2001	CH CH <sub>3</sub>
XA2002	CIH CH CH, CH, CH,
XA2003	
XA2004	HCI HCI N N CH <sub>3</sub>





XA2005	
, AA2003	HCI HCI CH <sub>3</sub>
XA2006	Ha Ha Ha N Ha CH <sub>3</sub>
XA2007	HCI HCI HCI HCI CH <sub>3</sub>





V. 2000	
XA2008	H <sub>3</sub> C S CH <sub>3</sub>
XA2009	HCI HCI CH <sub>3</sub>
XA2010	HCI HCI CH <sub>3</sub>
XA2011	N N N CH <sub>3</sub>





XA2012	H <sub>3</sub> C-SOOH NOCH <sub>3</sub>
XA2013	HCI HCI N HCI N CH <sub>3</sub>
XA2014	CH <sub>3</sub> HCI HCI N HCI N H HCI N HCI N H HCI N HCI N H HCI N H HCI N HCI N H HCI N H HCI N HCI N HC
XA2015	HO HO HO CH <sub>3</sub>





XA2016	HCI N HCI HCI CH <sub>3</sub>
XA2017	Ha Ha Ha Ha Ha CH <sub>3</sub>
XA2018	2-E-E-E-E-E-E-E-E-E-E-E-E-E-E-E-E-E-E-E
XA2019	H <sub>3</sub> C N N N N O CH <sub>3</sub>





VA2020	
XA2020	HO N N N N N N N N N N N N N N N N N N N
XA2021	H,C O O N N N N N N N N N N N N N N N N N
XA2022	2-2-2-2-2-2-3-3-3-3-3-3-3-3-3-3-3-3-3-3
XA2023	THE CONTRACTOR OF THE CONTRACT
XA2024	HO—\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\





XA2025	H <sub>3</sub> C N N N O
XA2026	CH <sub>3</sub>
	H <sub>3</sub> C N N N O CH <sub>3</sub>
XA2027	H <sup>2</sup> C S S S S S S S S S S S S S S S S S S S
XA2028	Z CH <sub>3</sub>





V40000	
XA2029	
XA2030	F CH <sub>3</sub>
XA2031	H <sub>3</sub> C
XA2032	N N CH <sub>3</sub>

V40000	
XA2033	H <sub>3</sub> C
XA2034	CH3 CH3 CH3 O
XA2035	CH <sub>3</sub> O CH <sub>3</sub>





VADOOD	
XA2036	
XA2037	
XA2038	CI CH <sub>3</sub>
XA2039	ON CH <sub>3</sub>

XA2040	N N N N N N N N N N N N N N N N N N N
XA2041	
XA2042	H <sub>3</sub> C O O O O O O O O O O O O O O O O O O O
XA2043	H <sub>2</sub> C \ 0 \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \





XA2044	
	CH <sub>3</sub> S O CH <sub>3</sub> O CH <sub>3</sub> O CH <sub>3</sub> O
XA2045	H <sub>3</sub> C N N N O CH <sub>3</sub>
XA2046	H <sub>3</sub> C CH <sub>3</sub> H <sub>3</sub> C CH <sub>3</sub> O CH <sub>3</sub>
XA2047	H <sub>3</sub> C N N N O CH <sub>3</sub>

V42048	T
XA2048	H <sub>2</sub> N CH <sub>3</sub>
XA2049	H <sub>3</sub> C N N N O CH <sub>3</sub>
XA2050	B Z Z Z Z Z Z Z Z Z Z Z Z Z Z Z Z Z Z Z
XA2051	Br Z-H <sub>3</sub>





140000	
XA2052	Br CH <sub>3</sub>
XA2053	nN
	H <sub>3</sub> C OH <sub>3</sub> O
XA2054	N
YAQQEE	H <sub>C</sub> C
XA2055	
	H <sub>3</sub> C





Table-2				•	
		R <sub>2</sub> N N N N N N N N N N N N N N N N N N N			
No	R1	R2	R3	R4	R5
XB1	снз-	СН3-	Н	Н	н .
XB2	СН3-	снзсн2-	н	Н	н
XB3	СН3-	<b>^</b> \	н	н	н
XB4	СН3-	Y	н	Н	н
XB5	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н	н
XB6	снз-	L	Н	н	н
XB7	снз-	7,	н	Н	н
XB8	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н.	Н	н
XB9	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Н	Н
XB10	СН3	<b>\\\\</b>	н	Н	н
XB11	СН3-	~~~``\	н	н .	н
XB12	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Н	н
XB13	СН3-		н	Н	н
XB14	СН3-		н	Н	н
XB15	СН3-		н	н .	н
XB16	снз-		н	н	Н
XB17	CH3	F	н	H.	н

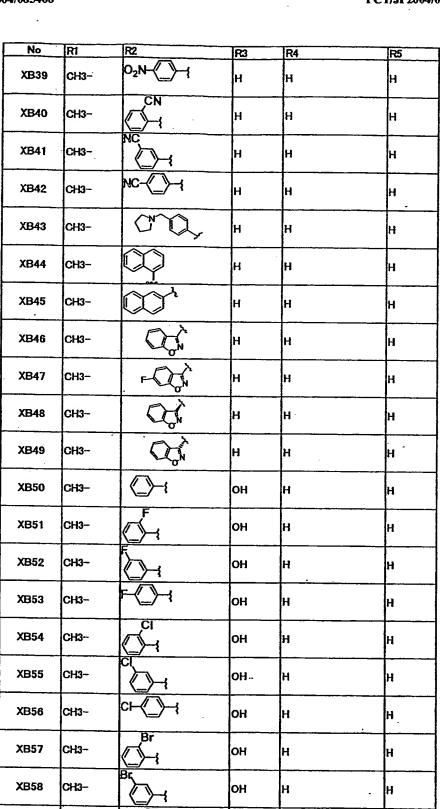


N.	104	Ipa	Ina	R4	R5
No	R1	R2	R3	174	
XB18	снз-		н	н	н
XB19	снз-		н	н	Н
XB20	снз-	CI	н	Н	н
XB21	снз-	Ci	Н	Н	Н .
XB22	СН3-	C⊢{}~{	н	н	н
XB23	снз	Br	н	н	н
XB24	снз-	Br.	Н	Н	H
XB25	снз-	Br—⟨	н	Н	н
XB26	CH3-	CH <sub>3</sub>	н	н	Н
XB27	снз-	H <sub>3</sub> C	н	н	Н
XB28	снз-	H <sub>3</sub> C-{}-{	н	н	Н
XB29	снз-	C <sub>2</sub> H <sub>5</sub> -{_}-{	. Н	н	H
· XB30	снз-	OH OH	н	Н	н
XB31	снз-	НО	Н	н	н
XB32	СН3-	HO-{_}-	н	н	н
XB33	СН3-	OCH₃	н	н	н
XB34	СН3-	H₃CO >(	н	н	н
XB35	СН3-	H <sub>3</sub> CO-{	н	н	н
XB36	СН3-	C <sub>2</sub> H <sub>5</sub> O-{}-{	н	н	н
XB37	снз-	NO <sub>2</sub>	н	Н	Н
XB38	снз-	O <sub>2</sub> N	н	н	н



XB59

CH3-



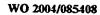
ОН

Н

Н



□ No	Tos	Ine	100	Ine	loc .
No	R1	CH <sub>3</sub>	R3	R4	R5
XB60	снз-		он	н	н
XB61	снз-	ӊс ҈҅	он	н	Н
XB62	снз-	H₃C-{_}	он	н	Н
XB63	снз-		он	Н	Н
XB64	снз-	OH OH	он	н	н
XB65	снз-	HO	он	Н	н
XB66	снз-	HO-{\(\){\}	он	н	н
XB67	снз-	осн₃	он	Н	н
XB68	снз-	H <sub>3</sub> CO	он	н	Н
XB69	СН3	н₃со-⟨∑⊢;	он	н	H'
XB70	снз-	C <sub>2</sub> H <sub>5</sub> O-{	он	н	Н
XB71	снз-	NO <sub>2</sub>	он	н	н
XB72	снз	O <sub>2</sub> N	он	н	н
XB73	СН3-	O <sub>2</sub> N-{_}	он .	н	н
XB74	снз-	CN ◯→	он	н	н
XB75	снз-	NC	он	н	н
XB76	снз-	NC-{\rightarrow}-{	он .	н	н
XB77	СН3-	ara,	он	н	н
XB78	снз-		он	Н	Н
XB79	снз-	CCT'	ОН	н	н
XB80	снз-		CN	н .	Н



No	- R1	R2	R3	ID4	lng.
XB81	СН3-	F	CN	R4 H	R5
XB82	снз-	F	CN	н	н
XB83	СН3-	F-()-1	CN	Н	н
XB84	СН3-	CI	CN	н	н
XB85	снз-	CI_	CN	н	н
XB86	снз-	CH	CN	н	н
XB87	снз-	Br	CN	н	н
XB88	СН3-	Br	CN	н	Н
XB89	снз-	Br-{\rightarrow}-{	CN	н	Н
XB90	снз-	CH₃	CN	н	н
XB91	СН3-	H <sub>3</sub> C	CN	н	н
XB92	СН3-	H <sub>3</sub> C-\_	CN	н	н
XB93	СН3-	$C_2H_5$	CN	Н	н
XB94	СН3-	○H OH	CN	н	н
XB95	СНЗ-	HO T	CN	н	н
XB96	СН3-	HO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar	CN	Н	Н
XB97	СН3-	OCH₃	CN	н	Н
XB98	снз-	H <sub>3</sub> CO	CN	н	Н
XB99	снз	H³CO- <b>⟨</b> }~-{	CN	Н	н
XB100	СН3-	C <sub>2</sub> H <sub>5</sub> O-{	CN	Н	Н
XB101	СН3-	NO <sub>2</sub>	CN	Н	н



No	R1	R2	R3	R4	R5
XB102	СН3-	O <sub>2</sub> N	CN	Н	н
XB103	снз-	O <sub>2</sub> N-{	CN	н	н
XB104	снз-	CN C	CN	Н	Н
XB105	снз-	NC	CN	Н	н
XB106	снз-	NC-{}	CN	Н	Н
XB107	снз-	aro,	CN	Н	н
XB108	снз-		CN	н .	Н
XB109	СН3-	CC,	CN	н	Н
XB110	снз-	н	Н	СН3-	H
XB111	снз-	н	н	СНЗСН2-	Н
XB112	снз-	н	н	<b>^</b> \	н
XB113	снз-	н	н	<b>Y</b>	н
XB114	снз-	н	н	<b>~~</b> `\	H
XB115	снз-	Н	н	人人	Н
XB116	снз-	H .	н	<b>Y</b>	Н
XB117	снз-	н	н	~~``	н
XB118	снз-	н	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н
XB119	снз-	н	н	<b>~~</b>	н
XB120	снз-	н	н	~~~`\	н
XB121	снз–	н	Н	<b>~~~</b>	H
XB122	СН3-	Н	Н	Qu	н



No	R1	R2	Too .	154	lese.
	- I''	112	R3	R4	R5
XB123	снз-	H	н	U <sub>A</sub>	н
XB124	CH3	н	н	OCH,	н
XB125	снз-	н	Н		н
XB126	CH3-	н	н		Н .
XB127	снз-	Н	н		Н
XB128	снз-	H ·	н	F	Н
XB129	снз-	н	. н		н
XB130	снз-	H	н	F-()-{	Н
XB131	снз-	Н	Н	CI	н
XB132	снз-	н	н	CI	Н
XB133	СН3-	H .	н	c⊢(_)~{	Н
XB134	СН3-	н .	Н	ci ci—	н
XB135	СН3-	н	Н	Br	н
XB136	снз-	Н	н	Br.	н
XB137	СН3	н	н	Br-{_}-{	Н
XB138	снз-	Н	н	CH₃	н
XB139	снз-	Н	. Н	H₃C ———	н
XB140	снз-	Н	Н	H3C-{}-{	Н
XB141	СН3-	н	Н	C <sub>2</sub> H <sub>5</sub> -{}-{	н
XB142	снз-	н	н	OH ○	н
XB143	снз-	н	Н	HO	н



No	R1	R2	R3	R4	R5
	CH3-	H	H	HO-{}-{	н
XB145	CH3-	н	н	OCH₃	Н
XB146	CH3-	н	н	H <sub>3</sub> CO	н
XB147	снз-	н	н	H₃CO-{_}-{	Н .
XB148	СН3-	Н	Н	C <sub>2</sub> H <sub>5</sub> O-{	н
XB149	СН3-	Н	н	NO <sub>2</sub>	Н
XB150	СН3-	н	н	O <sub>2</sub> N	н
XB151	СН3-	н	н		Н
XB152	СН3-	н	н	CN	н
XB153	СН3-	н	н	NC.	н
XB154	СН3-	Н	н	NC-{}-{	н
XB155	СН3-	Н	Н		н
- XB156	СН3	н	н	CO,	н
XB157	СН3-	н	н	FOX	н
XB158	СН3-	н	н	FON	Н
XB159	СН3-	н	н	FOR	н
XB160	СН3-	н	н .		н
XB161	СН3-	Н	Н	CXN	н
XB162	СН3-	Н	н	<b>○</b>	н
XB163	СН3-	Н	н	Oh,	н
XB164	снз-	н	н	Oh	Н





No	R1	R2	R3	R4	R5
XB165	СН3-	н	н	CH <sub>3</sub>	Н
XB166	снз-	н	Н	F N CH <sub>3</sub>	Н
XB167	снз	н	Н	H <sub>3</sub> C O	Н
XB168	СН3-	н	Н	H <sub>3</sub> CO	н

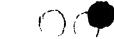




No	R1	R2	R3	Ine	- Inc
		<del></del>	<u> ns</u>	R4	R5
XB169	снз-	Н	н		он
XB170	СН3-	н	Н	F	он
XB171	СН3-	н	Н	F.	он
XB172	СН3-	н	н	F-()1	он
XB173	СН3-	н	. Н	CI	он
XB174	СН3-	н	н	CI	он
XB175	снз-	Н	Н	C <b>├</b> ──}	он
XB176	снз-	н	н	Br {	он
XB177	снз-	н	Н	Br.	он
XB178	снз-	н	н	Br—{}	он
XB179	снз	н	н	CH₃	он
XB180	снз-	н	Н	H <sub>3</sub> C	он
XB181	СН3-	н	н	H <sub>3</sub> C-{}-{	он
XB182	снз-	н	Н	C <sub>2</sub> H <sub>5</sub> —{	он
XB183	снз-	н	н	OH OH	он
XB184	СН3-	н	Н	HO	он
XB185	снз–	н	Н	HO-{}-{	он



No	R1	R2	lR3	R4	R5
XB186	снз-	н	н	OCH₃	он
XB187	снз-	н	н	н₃со	он
XB188	снз-	н	н	H <sub>3</sub> CO-{}-{	он
XB189	СН3-	н	н	C <sub>2</sub> H <sub>5</sub> O-{	он .
XB190	снз-	н	н	NO <sub>2</sub>	он
XB191	снз-	Н	Н	O <sub>2</sub> N	он
XB192	снз-	Н	Н	O <sub>2</sub> N-{}-{	он
XB193	снз-	Н	Н	CN	он
XB194	снз-	н	н	NC	ОН
XB195	снз-	н	н	NC-{\rightarrow}	он
XB196	СН3-	н	н	CQ	он
XB197	снз-	н	н	CC,	он
XB198	снз-	н	н		CN
XB199	снз-	Н	н		CN
XB200	СН3-	Н	н		CN
XB201	СН3-	н	н	F-(){	CN
XB202	снз-	н	н	CI	CN
XB203	снз-	н	Н	CI.	CN .
XB204	снз-	Н	н	c⊢(_ <del>\</del>	CN
XB205	снз-	Н	н	Br	CN
XB206	СН3-	н	Н	Br.	CN



No	RI	R2	R3	R4	R5
	1		1100	43	rto .
XB207	CH3-	н	H	Br- <b>⟨_</b> )→{	CN
XB208	снз-	н .	н	CH <sub>3</sub>	CN
XB209	СН3-	н	н	H <sub>3</sub> C	CN
XB210	снз-	н	н	H₃C- <b>(</b> ){	CN
XB211	СН3-	н	Н	C <sub>2</sub> H <sub>5</sub> -{}-{	CN
XB212	СН3-	н	Н	OH OH	CN
XB213	СН3-	н	н	HO —	CN
XB214	снз-	н	Н	HO-{	CN
XB215	СН3-	н	н	OCH <sub>3</sub>	CN
XB216	СН3-	н	н	H <sub>3</sub> CO	CN
XB217	снз-	н	н	H³CO-{_}-4	CN
XB218	CH3-	н	H	C <sub>2</sub> H <sub>5</sub> O-{_}-	CN
XB219	снз-	н	Н	NO <sub>2</sub>	CN
XB220	снз-	н	. н	O <sub>2</sub> N	CN
. XB221	СН3-	н	н	O <sub>2</sub> N-{}	CN
XB222	СН3-	н	н	CN	CN
XB223	снз-	н	н.	NC	CN
XB224	снз-	н	н	NC-{}	CN
XB225	снз-	Н	н		CN
XB226	снз-	Н	н	CC,	CN
XB227	снз-	Н	н		<u>}</u> ,





No	R1	R2	R3	R4	R5
XB228	снз-	Н	Н	F	<u></u>
XB229	снз	н	н	F	Ů,
XB230	снз-	н	н	F-()-1	2,
XB231	СН3	н	н	CI C)	· ·
XB232	СН3-	н	Н	CI	9
XB233	снз-	н	Н	CH(_ <del>}</del> →	0
XB234	СН3-	н	н	Br	<u></u>
XB235	СН3-	н .	Н	Br.	0
XB236	снз-	н	Н	Br- <b>√</b>	<u></u>
XB237	снз–	н	н	CH₃	<u>\</u>
XB238	снз-	н	н	H₃C ————————————————————————————————————	<u></u>
XB239	СН3-	н	н	H³C-⟨_}~	<u></u>
XB240	снз-	н	Н		<u>}</u>
XB241	снз-	н	н	OH.	<u>}</u> ,
XB242	СН3-	н	н	HO —	بُ
XB243	СН3-	н	Н	HO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar	3,
XB244	СН3	н	н	OCH₃	3
XB245	снз-	н	н	H₃CO ☐	Ĵ,
XB246	снз-	Н	н	H <sub>3</sub> CO-{}-{	Î,
XB247	снз-	н	н	C <sub>2</sub> H <sub>5</sub> O-{	2
XB248	снз-	н	н	NO <sub>2</sub>	0





No	R1	R2	P3	R4	R5
XB249	снз-	н	н	O <sub>2</sub> N	0
XB250	снз-	н	н	O <sub>2</sub> N-{}	<u></u>
XB251	снз-	н	н	CN	<u></u>
XB252	снз-	Н	Н	NC	°,
XB253	СН3-	н	H	NC-{}	<u></u>
XB254	снз-	Н	Н		<u></u>
XB255	снз-	н	н.	CC,	Ŷ,



No.	STRUCTURE
	·
XB256	
	N N N O CH <sub>3</sub>
XB257	N. M.
L	CH <sub>3</sub>
XB258	·
	CIH N N O CH <sub>3</sub>
XB259	N
	CH <sub>3</sub>

XB260	
•	CIH N N O O O O O O O O O O O O O O O O O
XB261	N CH <sub>3</sub>
XB262	H <sub>3</sub> C N N N O CH <sub>3</sub>
XB263	CIH CIH CIH N N N N O CH <sub>3</sub> CH <sub>3</sub>

VD0C4	
XB264	N N
	H <sub>3</sub> C N CH <sub>3</sub>
XB265	N
VP000	H <sub>3</sub> C N N N O CH <sub>3</sub>
XB266	
	ан Си
VDOOZ	H <sub>3</sub> C-N <sub>CH<sub>3</sub></sub> CIH
XB267	N
	Br CH <sub>3</sub>
XB268	
	Br CH,





XB269	
XB270	- Ct.
XB271	F CH <sub>3</sub>
XB272	F F N N O CH <sub>3</sub>





XB273	H <sub>3</sub> C N N N O CH <sub>3</sub>
XB274	CH <sub>3</sub> N CH <sub>3</sub> O C CH <sub>3</sub> O C CH <sub>3</sub> O C CH <sub>3</sub> O C C C C C C C C C C C C C C C C C C
XB275	CH <sub>3</sub>
XB276	CH <sub>3</sub> N O CH <sub>3</sub>





XB277	T The second sec
XB278	CH <sub>3</sub>
XB279	CH <sub>2</sub> CH <sub>3</sub>
XB280	H <sub>3</sub> C CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub>
XB281	Br N N O CH <sub>3</sub>



VDOOD	
XB282	O CH3
XB283	HO N CH <sub>3</sub>
XB284	H <sub>3</sub> C N O CH <sub>3</sub>
XB285	ON CH3
XB286	ON CH <sub>3</sub>





XB2 <u>8</u> 7	H <sub>3</sub> C N N N N O CH <sub>3</sub>
XB288	H <sub>3</sub> C N N N N N CH <sub>3</sub>
XB289	3-2-2
XB290	H,C N





XB291	
	HO CH <sub>3</sub>
XB292	N
-	N N N O CH <sub>3</sub>
XB293	
YP204	CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub>
XB294	H,C OCH OCH OCH OCH OCH OCH OCH OCH OCH OC
XB295	CH <sub>3</sub> CH <sub>3</sub> CH <sub>3</sub> O





	<u> </u>
XB296	CH3 CH3
XB297	H <sub>3</sub> C OH <sub>3</sub>
XB298	H <sub>3</sub> C
XB299	N CH <sub>3</sub>





VDago	
XB300	CH <sub>3</sub>
XB301	ON CH <sub>3</sub>
XB302	CH <sub>3</sub>



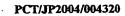


Table-3				
		<b>X</b> "		
		R <sup>2</sup> R <sup>2</sup>		
1		R N N N		
No.	R1			
YA0001	CH3-	R2 H	R3 H	R4 CH3-
YA0002	СН3-	Н	H	CH3CH2-
YA0003	СН3-	Н	Н	<b>△</b> \\
YA0004	СН3-	Н	Н	Y
YA0005	СН3	Н	н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\
YA0006	CH3-	Н	н	人心
YA0007	СН3	Н	Н	Y
YA0008	CH3-	Н	н	Y
YA0009	СН3-	н	н	Qu
YA0010	· CH3-	н	н	
YA0011	СН3-	Н	н	Q.
YA0012	СН3-	. н	н	$\triangleright$
YA0013	снз-	н	н	$\Diamond$ -1
YA0014	CH3-	н	Н	$\bigcirc$
YA0015	CH3-	н	н	$\bigcirc$ -1
YA0016	СН3-	н	н	$\bigcirc$
YA0017	СН3-	Н	Н	
YA0018	СН3-	Н	Н	
YA0019	CH3-	Н	н	<u></u>
YA0020	СН3-	Н	н	F-()-
YA0021	CH3-	н	н	CI C)





No.	R1	R2	R3	T
		1/2	<del>  ~~</del>	R4
YA0022	CH3-	Н	Н	
YA0023	СН3-	н	Н	c <del>{-}-</del> {
YA0024	снз-	н	Н	Br
YA0025	СН3-	н .	н	Bt
YA0026	: CH3-	н	н	Br- <b>⟨)</b> →{
YA0027	СН3-	н	H	
YA0028	СН3-	н	н	
YA0029	СН3-	н	Н	
YA0030	СН3-	Н	Н	CH₃
YA0031	СН3-	Н	н	H₃C <u></u>
YA0032	СН3-	Н	н	H₃C- <b>(_)</b> —{
YA0033	снз-	Н	Н	C <sub>2</sub> H <sub>5</sub> -{}-{
YA0034	СН3-	Н	Н	n-C₃H₁−⟨∑⟩–{
YA0035	СН3-	H ·	Н	n-C <sub>4</sub> H <sub>9</sub> {}{
YA0036	СН3-	Н	н.	· OH
YA0037	CH3-	Н	Н	HO \
YA0038	CH3-	н	. Н	HO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar
YA0039	СН3-	Н	Н	OCH₃
YA0040	СН3-	н	н	H₃CO <b>(_</b> )~-
YA0041	СН3-	н	Н	н₃со-{_}-{
YA0042	СН3-	Н	н	C <sub>2</sub> H <sub>5</sub> O-{}-{







No.	R1	R2	R3	R4
YA0043	снз-	н	Н	n-C <sub>3</sub> H <sub>7</sub> O-{}{
YA0044	CH3	н	Н	n-C <sub>4</sub> H <sub>9</sub> O-
YA0045	CH3-	н	н	NO <sub>2</sub>
YA0046	снз-	н	Н	O <sub>2</sub> N
YA0047	снз-	н	Н	O <sub>2</sub> N-{_}
YA0048	снз-	Н	н	CN CN
YA0049	СН3-	Н	н	NC
YA0050	СН3-	н .	н	NC-(
YA0051	СН3-	Н	н	CF <sub>3</sub>
YA0052	СН3-	н	н	F <sub>3</sub> C
YA0053	CH3-	Н	.H	F <sub>3</sub> C-{\_}-{
YA0054	СН3-	Н	н	COOH COOH
YA0055	СН3-	Н	н	HOOC
YA0056	СН3	н	н	HOOC-{\(\){\}
YA0057	СН3-	н.	н	CO <sub>2</sub> Me
YA0058	CH3-	н	Н	MeO <sub>2</sub> C
YA0059	CH3~	н	Н	MeO <sub>2</sub> C-⟨}
YA0060	CH3-	Н	Н	CO <sub>2</sub> Et
YA0061	CH3-	н	н	EtO <sub>2</sub> C
YA0062	CH3-	н	н	EtO <sub>2</sub> C-{}
YA0063	CH3-	н	н	SMe ∠;



<del></del>			700	R4
No.	R1	R2	R3	
YA0064	СН3-	н	н	MeS
YA0065	СН3-	Н	н	MeS-{_}-{
YA0066	СН3-	н	н	SO₂Me
YA0067	СН3-	Н	Н	MeO <sub>2</sub> S
YA0068	снз-	н	Н	MeO <sub>2</sub> S-{
YA0069	CH3-	н	Н	NH <sub>2</sub>
YA0070	СН3-	Н	н	H <sub>2</sub> N
YA0071	CH3-	н	н	H₂N-{\rightarrow}-{\rightarro
YA0072	СН3-	н	н	NMe₂
YA0073	СН3	Н	н	Me <sub>2</sub> N
YA0074	CH3-	н	н	Me₂N-⟨¯){
YA0075	снз-	н	н	
YA0076	СН3-	н	н	OU'
YA0077	снз-	н	н	C)
YA0078	снз-	н	н	O'N
YA0079	CH3-	н	н	
YA0080	снз-	н	н	Fi,





No.	R1	R2	R3	R4
YA0081	CH3-	н	н	F
YA0082	CH3-	н	Н	F.O.
YA0083	снз-	Н	н	G o
YA0084	СН3-	Н	Н	a گړ <sup>ې</sup>





				• •
No.	R1	R2	R3	R4
YA0085	снз-	н	Н	a Cir
YA0086	снз-	H	Н	Br Q
YA0087	снз-	H	Н	Br
YA0088	СН3-	н	н	Br
YA0089	CH3-	н	н	CHO
YA0090	CH3-	н	Н	H₃C C
YA0091	снз-	н	н	H <sub>3</sub> C , ,
YA0092	CH3-	Н	н	CH3O O
YA0093	CH3	Н	н	H <sub>3</sub> CO
YA0094	СН3-	Н	Н	H <sub>3</sub> CO
YA0095	СН3-	Н	н	NO.O
YA0096	CH3-	Н	н	O <sub>2</sub> N
YA0097	снз-	Н	Н	02N
YA0098	CH3-	Н	н	OH O
YA0099	СН3-	н	Н	но
YA0100	снз-	н	Н	но
YA0101	снз-	н	н	NH O





No.	R1	R2	R3	R4
YA0102	CH3~	Н	н	H <sub>2</sub> N
YA0103	CH3-	н	Н	H <sup>2</sup> M
YA0104	СН3-	Н	Н	CHO
YA0105	снз-	Н	Н	NC J.





			•	
No.	R1	R2	R3	R4
YA0106	CH3~	Н	н	NC ()
YA0107	СН3-	н	н	gi,
YA0108	СН3-	н	н	OO',
YA0109	CH3~	н	н	<u>ک</u> ہ
YA0110	CH3-	н	Н	<b>,</b>
YA0111	СН3-	н	Н	~ ,
YA0112	СН3-	н	Н	\ <sup>2</sup> ,
YA0113	СН3-	Н	Н	A.P.
YA0114	СН3-	Н	н	~~\ <sup>1</sup> ,
YA0115	CH3-	н	н	<del>}</del> }
YA0116	CH3-	Н	Н	~~ <sup>1</sup> ,
YA0117	СН3-	Н	H.	~~~ <sup>1</sup> ,
YA0118	СН3-	н	Н	~~~ <sup>Q</sup> ,
YA0119	СН3-	Н	Н	
YA0120	СН3-	Н	н	17°,
YA0121	СН3-	н	н	O'S'
YA0122	СН3-	Н	н	رائر





No.	R1	R2	R3	R4
YA0123	СН3-	O H³CO_≻	Н	Н
YA0124	CH3-	H³CO_≻	Н	снз-
YA0125	CH3-	H³CO_>	Н	СН3СН2-
YA0126	CH3-	H³CO_} N	Н	<b>∕</b> ∕\





No.	R1	R2	R3	R4
<del></del>			<del>                                     </del>	人 入 "**
YA0127	CH3-	H³CO, ≻	н	Υ,
YA0128	СН3-	H³CO ≻	Н	<b>~</b> ~
YA0129	СН3-	о н₃co <sup>*</sup> ≻	Н	人、
YA0130	CH3-	H³CO, ∆	н	~
YA0131	CH3-	о Н <sub>3</sub> со <sup>7</sup> У	Н	丫
YA0132	CH3-	H³CO, ≻	н	
YA0133	CH3-	H³CO,≻	н	
YA0134	снз-	H³CO, ≻	н	
YA0135	СН3~	H³CO_≻	н	<b>⊳</b> ⊣
YA0136	CH3-	H₃CO →	Н	$\Diamond$ -1
YA0137	CH3-	H³CO, ≻	н	$\bigcirc$
YA0138	CH3-	H³CO_≻	Н	$\bigcirc$ $\dashv$
YA0139	СН3-	H³CO,≻	н	
YA0140	CH3-	H³CO,≻	н	
ÝA0141	снз-	H³co_}	Н	
YA0142	, снз-	H³CO, ≻	н	
YA0143	снз-	H³CO,≻	Н -	F-()-1
YA0144	снз-	H³CO_≻ O	н	CI CI
YA0145	снз-	H³CO,≻	н	
YA0146	снз-	H <sub>3</sub> CO >	Н	c⊢
YA0147	CH3-	H³CO,≻	Н	Br ∰





No.	D1			
	R1	R2	R3	R4 Br,
YA0148	CH3-	H3CO,	н	
YA0149	СН3-	H₃CO →	н	Br-{}-{
YA0150	СН3-	Q H₃CO ≻	Н	CH <sub>3</sub>
YA0151	снз-	H³CO_>	Н	H <sub>3</sub> C
YA0152	СН3-	H₃CO >	н	H <sub>3</sub> C-{}-{
YA0153	CH3-	H₃CO >	н	C <sub>2</sub> H <sub>5</sub> —{
YA0154	снз-	H₃CO '>	н	n-C <sub>3</sub> H <sub>7</sub> -
YA0155	СН3-	H³CO, ≻	н	n-C <sub>4</sub> H <sub>9</sub> -
YA0156	CH3-	O H₃CO ≻	н	OCH <sub>3</sub>
YA0157	CH3-	H³CO, ³	н	H₃CO ⟨_)
YA0158	снз-	H <sub>3</sub> CO <sup>2</sup> >	н	H <sub>3</sub> CO-{\rightarrow}-{\rightarrow}-{\rightarrow}
YA0159	CH3-	H <sub>3</sub> CO 7	н	C <sub>2</sub> H <sub>5</sub> O-{
YA0160	снз-	H³CO, >	Н	n-C <sub>3</sub> H <sub>7</sub> O-
YA0161	снз-	H³CO, ≻ O	н	n-C₄H <sub>9</sub> O-{_}
YA0162	CH3-	O H₃CO ≻	H.	NO <sub>2</sub>
YA0163	CH3-	O H₃CO Y	н	O <sub>2</sub> N
YA0164	CH3-	о н <sub>з</sub> со <sup>д</sup> у	н	O <sub>2</sub> N-{}
YA0165	СН3-	H³CO_} →	Н	CN
YA0166	CH3-	H³CO_^≻ O	H	NC \_\
YA0167	СН3-	н <sub>з</sub> со <sup>т</sup> у	н	MC-{\rightarrow}-1
YA0168	СН3-	H <sub>3</sub> CO <sup>+</sup> >	н	NMe <sub>2</sub>



No.	R1	R2	R3	R4
YA0169	СН3-	O H₃CO ⁄	Н	Me <sub>2</sub> N
YA0170	CH3-	H³CO_≻	н	Me <sub>2</sub> N-{}
YA0171	СН3	O H₃CO →	Н	<u>Q</u>
YA0172	снз-	O H₃CO ≻	н	CO'r
YA0173	СН3-	H³CQ_≻	н	O <sup>l</sup> ,
YA0174	СН3-	H³CO_≻	н	Qi,
YA0175	CH3-	O H₃CO ≻	н	
YA0176	CH3	O H₃CO ≻	н	2,
YA0177	СН3-	H³CQ_≻ Ö	Н	<u></u>
YA0178	СН3	O C₂H₅O →	Н	Н
YA0179	СН3-	O C₂H₅O ∕≻	н	снз-
YA0180	СН3 <b>-</b>	C₂H₅O ≻	Н	СНЗСН2-
YA0181	СН3-	O C₂H₅O <sup>™</sup> ≻	н	<b>^</b> ∖\
YA0182	СН3-	C₂H₅O <sup>™</sup> ≻	н	Y
YA0183	СН3-	O C₂H₅O <sup>™</sup> ≻	Н	<b>\\\\\</b>
YA0184	CH3-	O C₂H₅O <sup>†</sup> >	н	





No.	R1	R2 -	R3	R4
YA0185	СН3-	O C₂H₅O 7′	Н	7
YA0186	CH3-	C⁵H²O , \	н	个
YA0187	CH3-	O C₂H₅O У	Н	Qu
YA0188	CH3-	O C₂H₅O →	н	
YA0189	СН3-	C₂H₅O У	н	





No	T 81			
No.	R1	R2	R3	R4
YA0190	CH3-	O C₂H₅O ≻	н	D-1
YA0191	CH3-	C <sub>2</sub> H <sub>5</sub> O >	н	$\Diamond$
YA0192	снз-	O C₂H₅O <sup>™</sup> ≻	Н	$\bigcirc$ -1
YA0193	СН3-	O C₂H₅O →	Н	$\bigcirc$ $\dashv$
YA0194	снз-	O C <sub>2</sub> H <sub>5</sub> O	н	OH
YA0195	снз-	O C₂H₅O y	Н	
YA0196	снз-	C₂H₅O √	н	
YA0197	СН3-	O C₂H₅O У	н	
YA0198	СН3-	O C <sub>2</sub> H <sub>5</sub> O /	H	
YA0199	СН3-	O C₂H₅O →	н	
YA0200	СН3-	O C <sub>2</sub> H <sub>5</sub> O >	н	
YA0201	снз-	O C₂H₅O <sup>†</sup> >⁄	н	cr()~;
YA0202	СН3-	O C <sub>2</sub> H <sub>5</sub> O	н	Br →
YA0203	снз-	O C <sub>2</sub> H <sub>5</sub> O /	н	Br.
YA0204	снз-	O C <sub>2</sub> H <sub>5</sub> O <sup>†</sup> >	H	Br—{_}_{
YA0205	снз-	O C <sub>2</sub> H <sub>5</sub> O ×	н	CH₃
YA0206	СН3-	O C₂H₅O У	Н	H₃C ⟨_}~
YA0207	снз-	O C₂H₅O <sup>™</sup> ≻	н	H <sub>3</sub> C-{_}-{
YA0208	CH3-	O C₂H₅O ∕ ≻	н	C2H5-{}-{
YA0209	СН3-	O C₂H₅O ≻	н	n-C <sub>3</sub> H <sub>7</sub> {}{
YA0210 -	CH3-	O C₂H₅O →	н	ı-C₄H <sub>9</sub> -⟨_}-{





	- B.		T 50	
No.	R1	R2	R3	R4
YA0211	СН3-	O C₂H₅O →	Н	OCH₃
YA0212	снз-	O C₂H₅O У	н	H₃CQ ————————————————————————————————————
YA0213	снз-	O C₂H₅O ✓	н	H₃CO-{_}-{
YA0214	СН3-	O C₂H₅O ✓	н	C <sub>2</sub> H <sub>5</sub> O-{
YA0215	CH3-	O C₂H₅O r	н	n-C₃H <sub>7</sub> O-{_}-{
YA0216	CH3-	Q C₂H₅O ≻	Н	n-C <sub>4</sub> H <sub>9</sub> O-
YA0217	СН3-	O C₂H₅O ✓	н	NO <sub>2</sub>
YA0218	СН3	O C₂H₅O ≻	Н	O <sub>2</sub> N
YA0219	СН3-	O C₂H₅O ≻	н	O <sub>2</sub> N-{}
YA0220	СН3-	O C₂H₅O →	Н	CN ←
YA0221	СН3-	O C₂H₅O r	н	NC.
YA0222	СН3-	O C₂H₅O <sup>™</sup> ⁄′	Н	NC-
YA0223	СН3-	O C₂H₅O <sup>^</sup> -⁄-	н	NMe <sub>2</sub>
YA0224	СН3-	O C₂H₅O <sup>™</sup> ≻	Н	Me <sub>2</sub> N
YA0225	снз-	C <sub>2</sub> H <sub>5</sub> O <sup>^</sup> , <sub>7</sub>	н	Me₂N-⟨\$\]
YA0226	снэ-	O C <sub>2</sub> H <sub>5</sub> O 7	н	





No.	R1	R2	R3	R4
· YA0227	CH3-	O C₂H₅O →	н	
YA0228	СН3-	O C₂H₅O →	Н	
YA0229	CH3-	O C₂H₅O →	н	
YA0230	СН3	O C <sub>2</sub> H <sub>5</sub> O >/	н	oo',
YA0231	СН3-	O C₂H₅O ∕≻	н .	<u>ک</u> ہ





No.	R1	R2	R3	R4
YA0232	CH3-	O C₂H₅O →	Н	Ŷ,
YA0233	снз-	CH3-	н	Н
YA0234	СН3-	СНЗСН2-	н	н
YA0235	СН3	<b>~</b> \	н	н
YA0236	CH3-	Y	н	н
YA0237	снз-	<b>~~</b> `\	н	н
YA0238	CH3-	人、	н	Н
YA0239	СН3-	~~	н	н
YA0240	снз-	丫	Н	Н
YA0241	СН3	~~``\	н	H
YA0242	CH3-	<b>/</b> ~	Н	н
YA0243	СН3-	X.	н	. н
YA0244	CH3-	*	. н	н
YA0245	СН3-	<b>&gt;&gt;&gt;</b>	н	н
YA0246	СН3-	<u>ل</u> مرح	н	Н
YA0247	СН3-	<b>^</b>	н	Н
YA0248	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
YA0249	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	н
YA0250	СН3-	L	н	н
YA0251	СН3-		Н	н
YA0252	снз-		Н	н



No.	R1	R2		·
YA0253	снз-	Own	R3 H	R4 H
YA0254	СН3-	D-1	н	н
YA0255	СН3-	$\Diamond$ -1	н	н
YA0256	СН3-	$\bigcirc$	н	Н
YA0257	CH3-	$\bigcirc$ $\dashv$	Н	Н
YA0258	СН3-	$\bigcirc$ $\dashv$	Н	н
YA0259	CH3-		Н	Н
YA0260	СН3		н	н
YA0261	снз-	<b></b>	н	Н
YA0262	СН3-		Н	Н
YA0263	СН3-		н	н
YA0264	СН3-	F-(){	н	н
YA0265	СН3-	F-()-(	Н	
YA0266	CH3-	F—(	н	н
YA0267	CH3-	CI	н	н
YA0268	CH3-		н	н
YA0269	снз-	CH	. н	Н
YA0270	СН3	CH	н	Н
YA0271	снз-		н	н
YA0272	снз-	Br	н	Н
YA0273	CH3-	35	Н	Н





<del></del>				
No.	R1	R2	R3	R4
YA0274	CH3-	Br—	н	Н
YA0275	снз-	Br—()	н	н
YA0276	CH3-	Br—(	н	н
YA0277	снз-		н	н
YA0278	снз-		н	Н
YA0279	СН3-		н	Н
YA0280	снз-	CH <sub>3</sub>	н	н
YA0281	CH3-	H₃C <u></u>	Н	Н
YA0282	CH3-	H <sub>3</sub> C-{_}-{	н	н
YA0283	СН3-	C <sub>2</sub> H <sub>5</sub> —{}	Н	н
YA0284	СН3-	n-C₃H <sub>7</sub> {_}}-{	н	Н
YA0285	CH3-	n-C₄H <sub>9</sub> ⟨}{	н	Н
YA0286	CH3-	OH OH	н	н
YA0287	CH3-	HO	н	Н
YA0288	СН3-	HO-{}-{	н	Н
YA0289	CH3	OCH₃	н	н
YA0290	снз-	H <sub>3</sub> CO	. Н	н
YA0291	СН3-	H₃CO <b>-</b> {}	н	Н
YA0292	CH3-	H₃CO- <b>(</b> )~{	н	н
YA0293	CH3-	H₃CO-⟨⟩⊪{	н	н
YA0294	снз-	OC <sub>2</sub> H <sub>5</sub>	н	н







No.	R1		· · · · · · · · · · · · · · · · · · ·	
740.	RI	R2	R3	R4
YA0295	CH3-		н	Н
YA0296	СН3-	C <sub>2</sub> H <sub>5</sub> O-⟨}-{	Н	. н
YA0297	снз-	n-C₃H <sub>7</sub> O-{_}-{	н	Н
YA0298	СН3-	n-C₄H₃O-{_}-{	н	н
YA0299	СН3-	NO <sub>2</sub>	Н	Н
YA0300	снз-	O <sub>2</sub> N	Н	н
YA0301	CH3-	O <sub>2</sub> N-{}	н	н
YA0302	снз-	CN ←	Н	н
YA0303	снз-	NC	н	н
YA0304	СН3-	NC-{\rightarrow}	Н	н
YA0305	СН3-	CF₃	Н	н
YA0306	CH3-	F <sub>3</sub> C	н	H
YA0307	снз-	F <sub>3</sub> C-{_}	н .	Н
YA0308	снз-	COOH	Н	Н
YA0309	CH3-	HOOC 	н	H
YA0310	CH3-	H00C-{\_}-{	н	н
YA0311	снз-	CO <sub>2</sub> Me	. н	н
YA0312	CH3	MeO₂C 	н	Н
YA0313	снз-	MeO₂C-⟨	н	Н
YA0314	СН3-	CO₂Et	н	н
YA0315	снз-	EtO <sub>2</sub> C	н	н



PCT/JP2004/004320



### WO 2004/085408

No.	R1	R2	R3	R4
YA0316	снз	EtO <sub>2</sub> C-	٠Н	н
YA0317	CH3-	SMe <_}-{	н	н
YA0318	CH3-	MeS	Н	н
YA0319	СН3~	MeS-{_}-{	Н	н
YA0320	снз-	SO₂Me	н	Н
YA0321	снз-	MeO <sub>2</sub> S	н	Н
YA0322	снз-	MeO <sub>2</sub> S-{_}	н	Н
YA0323	CH3-	NH <sub>2</sub>	н	Н
YA0324	СН3-	H <sub>2</sub> N →	н	н
YA0325	СН3-	H <sub>2</sub> N-{¯}-{	Н	Н
YA0326	СН3-	NMe <sub>2</sub>	н	Н
YA0327	CH3-	Me <sub>2</sub> N	н	Н
YA0328	СН3-	Me₂N-⟨¯}{	н	Н
YA0329	CH3-		н	Н
YA0330	СН3-		н	Н
YA0331	CH3-		н	Н
YA0332	CH3-	( )+ <u>( )</u>	н	Н
YA0333	CH3-	( )n-{_}}_	н	Н
YA0334	CH3-		Н	Н
YA0335	СН3-		н	Н
YA0336	CH3-	<b>○_N-</b> ()	н	н





No.	R1	R2	R3	R4
YA0337	СН3-	<b>€_N-{</b> _}-;	н	Н
· YA0338	снз-	H3CN_N-\	Н	Н
YA0339	CH3-	H₃CN N-⟨}	Н	Н
YA0340	снз-	H₃CN_N-{}-{	Н	н
YA0341	снз-	H₃C_CH₃	Н	н
YA0342	СН3-	H³C-{}-{	н	н
YA0343	СН3-	CH₃ H₃C	Н	Н
YA0344	снз-	CH <sub>3</sub>	Н	н
YA0345	снз-	H <sub>3</sub> C H <sub>3</sub> C-{}	Н	Н
YA0346	CH3-	H <sub>3</sub> C H <sub>3</sub> C	н	н
YA0347	снз-	F, F	н	н
YA0348	CH3-	F—F	н	н
YA0349	СН3-	\$\frac{F}{F}\$	н	н
YA0350	снз-	\ \times_F^F	н	н
YA0351	снз-	F——	н	н
YA0352	СН3-	F.	н	н





No.	R1	R2	R3	R4
YA0353	СН3-	a_a	н	Н
YA0354	снз-	a-√∑→	Н	н
YA0355	СН3-	a a	н	н
YA0356	СН3-	a∭a	н	н
YA0357	СН3-	a a————	н	н





No.	R1	R2	1 00	
····	NI	α κα	R3	R4
YA0358	CH3-		н	н
YA0359	СН3-	H3CO_OCH3	н	н
YA0360	снз-	H₃CO-{\bigsight\}	Н	, н
YA0361	СН3-	OCH₃ CD→ H₃CO	н	н
YA0362	CH3-	OCH <sub>3</sub>	н	н
YA0363	СН3-	H³CO-⟨}\}	н	н
YA0364	СН3-	H <sub>3</sub> CO	н	н
YA0365	снз-	F_OCH <sub>3</sub>	н	Н
YA0366	СН3-	OCH <sub>3</sub>	н	Н
YA0367	СН3-	OCH <sub>3</sub> F—⟨S	н	н
YA0368	СН3-	OCH <sub>3</sub>	н	н
YA0369	СН3-	OCH <sub>3</sub>	н	н.
YA0370	СН3-	OCH <sub>3</sub> F	н	н
YA0371	снз-	H₃CQ F-⟨□}	н	н
YA0372	СН3-	H <sub>2</sub> CQ F	Н.	н
YA0373	СН3-	H₃CO_F	н	н





No.	R1	R2	R3	R4
YA0374	СН3-	н₃со-{\$}	н	н
YA0375	CH3-	H <sub>3</sub> CO	н	н
YA0376	СН3-	H₃CO-{_}}	н	н
YA0377	СН3-	CI_OCH <sub>3</sub>	Н	н
YA0378	СН3-	OCH₃ CI—€	н	н

No.	R1	D2	l Do	D4
	<del>  ``</del>	R2 OCH <sub>3</sub>	R3	R4
YA0379	СН3-	a	н	н
YA0380	СН3-	G CCH3	н	н
YA0381	СН3-	H³CO	н	н
YA0382	снз-	H₃CQ CI	Н	н
YA0383	CH3-	H₃CO_CI ———;	н	Н
YA0384	снз-	H³co-{_}}-!	н	Н
YA0385	СН3-	, da , da , H₃co	н	н
YA0386	CH3-	CI, H₃CO-{}}	н	Н
YA0387	СН3-	F_CH <sub>3</sub>	н	Н
YA0388	СН3-	CH <sub>3</sub> F—	Н	н.
YA0389	CH3-	CH₃ F	н	н
YA0390	СН3-	EtH3	Н	H ×
YA0391	СН3-	H <sub>3</sub> C F—(	Н	н
YA0392	СН3-	H <sub>3</sub> C	Н	н
YA0393	СН3-	H₃C_F	Ħ	н
YA0394	СН3-	H₃C-{\(\)}	н	н





No.	R1	R2	R3	R4
YA0395	СН3-	H₃c H₃c	Н	н
YA0396	СН3-	H₃C-{}}-{	н	н
YA0397	СН3-	Br_OCH <sub>3</sub>	Н	H <sub>.</sub>
YA0398	СН3-	OCH₃ Br—∰	Н	н
YA0399	СН3-	OCH₃ SH	н	н





>				
No.	R1	R2	R3	R4
YA0400	СН3-	OCH <sub>3</sub>	н	н
YA0401	CH3-	H₃CO Br—{	н	н
YA0402	CH3-	H <sub>3</sub> CO Br	Н	н
YA0403	снз-	H <sub>3</sub> CO_Br	н	н
YA0404	снз-	H <sub>3</sub> CO-{  Sr	Н	н
YA0405	СН3-	Br H₃co	н	н
YA0406	СН3-	H <sub>3</sub> CO	н	н
YA0407	снз-	UN-{_} H³CO }	н	Н
YA0408	СН3-	CN-COCH3	Н	н
YA0409	снз-	Cu-⟨_}-ocH³	н	н
YA0410	СН3-	H3CO_}-N()	н	н
YA0411	СН3-	H³CO	н	н
YA0412	СН3-	OCH3	н	Н
YA0413	СН3-	F F	Н	н
YA0414	СН3-	OCH₃ F————————————————————————————————————	н	Н





No.	R1	R2	R3	R4
YA0415	СН3-	H₃CO-{\}; F	н	н
YA0416	СН3-	OCH <sub>3</sub>	н	н
YA0417	СН3-	0CH <sub>3</sub> H <sub>3</sub> CO-⟨_}_} OCH <sub>3</sub>	H	н
YA0418	СН3-	a a-{∑}, a	н	н
YA0419	СН3-	οα+₃ α-⟨∑-¦ α	Н	н
YA0420	СН3-	а н₃∞-{∑-; а	н	н



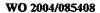


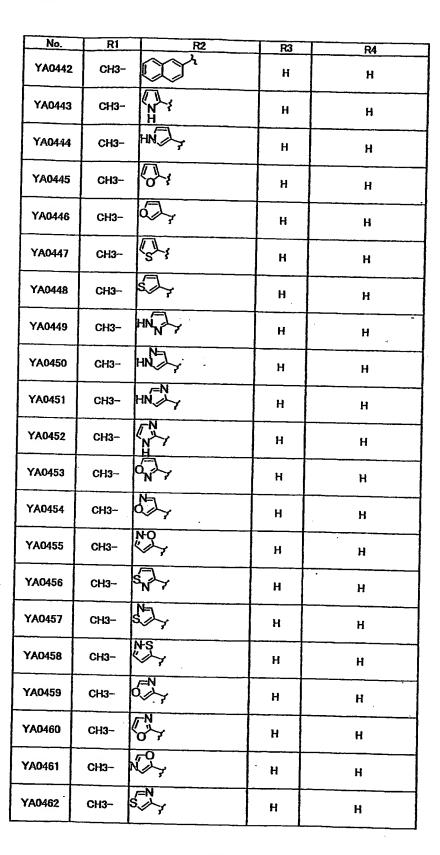
N-	1 ps			
No.	R1	R2 OCH <sub>3</sub>	R3	R4
YA0421	CH3-	CI(∑)-I	н	н
YA0422	снз-	H³CO-⟨_}-\ H°CO-⟨_}-\ OCH²	Н	Н
YA0423	снз-	OCH <sub>3</sub>	н	н
YA0424	снз-	H³CO	Н	н
YA0425	снз-	H₃CO-{\right\}-{\right\}-{\frac{1}{2}}-{\frac{1}{2}}	н	н
YA0426	снз-	OCH <sup>3</sup> }	н	Н
YA0427	СН3-	H <sub>3</sub> CO \	Н	H .
YA0428	снз–	H₃co-{	Н	н
YA0429	СН3	OCH <sub>3</sub>	н	н
YA0430	СН3-	H,CQ	н	н
YA0431	снэ–	н₃со-⟨҈\	Н	Н
YA0432	снз-	<b>₫</b>	Н	Н
YA0433	снз-	<b>E</b>	н	Н
YA0434	снз	F-(-)-(-)-1	н	н
YA0435	СН3-		Н	Н
YA0436	снз-		н	н





No.	R1	R2	R3	R4
YA0437	СН3-	F-()-()	н	н
YA0438	СН3-	4	Н	`н
YA0439	СН3-		н	н
YA0440	СН3-		н	н
YA0441	СН3-		Н	Н



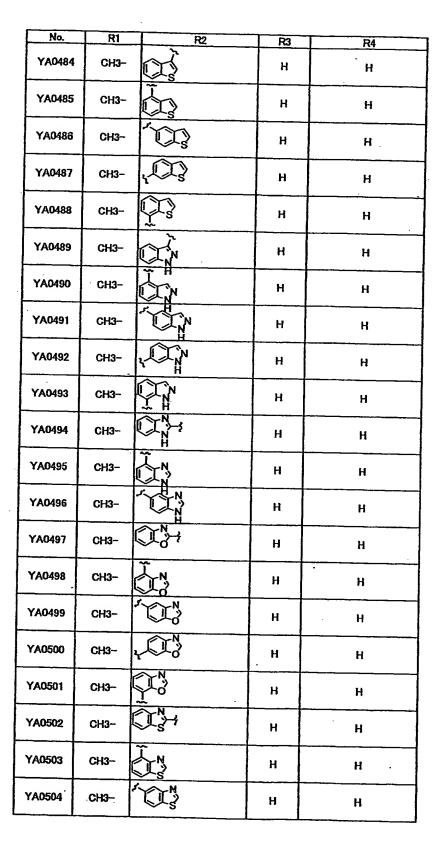


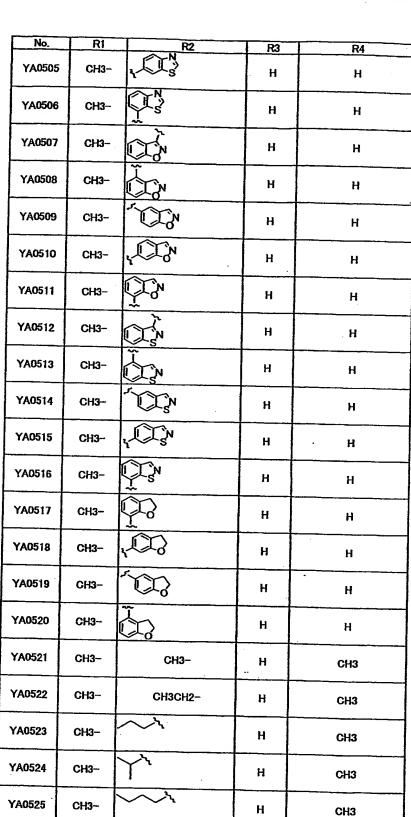


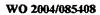
No.	R1	R2	R3	R4
YA0463		SN	Н	Н
YA0464	СН3-	√S	н	Н
YA0465	снз-		н	Н
YA0466	СН3-	N-	Н	Н
YA0467	СН3		Н	Н
YA0468	СН3-	CN-1	н	Н
YA0469	СН3-	NC)-1	Н	н
YA0470	СН3-	N-	н	н
YA0471	СН3-		Н	н
YA0472	CH3-		н	Н
YA0473	CH3-		н	Н
YA0474	снз-		н	Н
YA0475	CH3-	,Ch	н	н
YA0476	CH3-		Н	н '
YA0477	CH3-	CI}-1	Н	Н
YA0478	СН3-	CI)	н	н
YA0479	CH3-	Ü;	Н	н
YA0480	CH3-	(C)	н	н
YA0481	СН3-	,CI)	Н	Н
YA0482	СН3-	Ď3	н	. н
YA0483	снз-		н	н











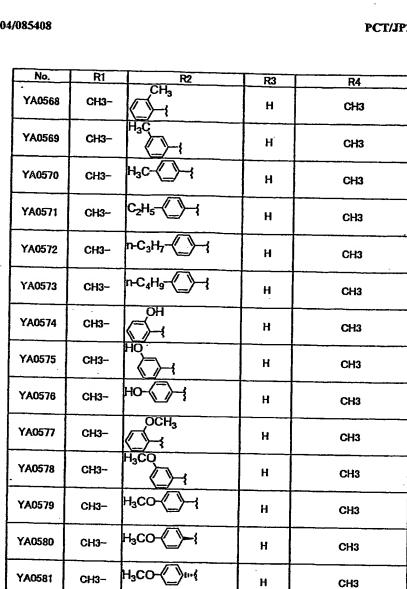


No.	R1	R2	1 00	
		1	R3	R4
YA0526	CH3-		Н	снз
YA0527	CH3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	СНЗ
YA0528	CH3-	丫	Н	СНЗ
YA0529	CH3-	~~\r\	Н	СНЗ
YA0530	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	снз
YA0531	СН3-	X.	н	снз
YA0532	снз-	7	Н	снз
YA0533	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	СНЗ
YA0534	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	снз
YA0535	снз-	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СНЗ
YA0536	снз-	~~~	н	СН3
YA0537	CH3-	<b>~~~</b>	Н	СНЗ
YA0538	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СНЗ
YA0539	СН3	Q	Н	СНЗ
YA0540	CH3-		Н	CH3
YA0541	СН3-		н	СНЗ
YA0542	CH3-	$\triangleright \dashv$	. н	СНЗ
YA0543	CH3-	$\Diamond$ -I	н	СНЗ
YA0544	снз-		н	СНЗ
YA0545	СН3-	$\bigcirc$ $\vdash$	н	СНЗ
YA0546	CH3-	$\bigcirc$ $\dashv$	н	СНЗ





	<del></del>	T		
No.	R1	R2	R3	. R4
YA0547	СН3-		н	СНЗ
YA0548	снз-		Н	СНЗ
YA0549	снз-	()mf	н	СНЗ
YA0550	СН3		н	СНЗ
YA0551	CH3-		н	СНЗ
YA0552	СН3-	F-()-1	Н	СНЗ
YA0553	снз-		Н	СНЗ
YA0554	CH3-	F-()  -(1)	н	СНЗ
YA0555	СН3-	CI CI	н	СНЗ
YA0556	СН3-	CI ————————————————————————————————————	, н	СНЗ
YA0557	СН3-	C⊢∕}	н	СНЗ
YA0558	CH3-	c⊢ <b>(_</b> }-	Н	СНЗ
YA0559	СН3-	CI()···(	н	СН3
YA0560	CH3-	Br	н	СНЗ
YA0561	СН3-	Br.	н	СНЗ
YA0562	СН3-	Br-{}	н	снз
YA0563	СН3-	Br-()	н	снз
YA0564	СН3-	Br—(	н	снз
YA0565	снз-		н	СНЗ
YA0566	СН3		н	CH3
YA0567	снз-	H	н	снз



OC<sub>2</sub>H<sub>5</sub>

'n-C₄H₀O-

NO<sub>2</sub>

H

Н

Н

Н

Н

Н

Н

СНЗ

СНЗ

СНЗ

CH3

СНЗ

СНЗ

СНЗ

YA0582

YA0583

YA0584

YA0585

YA0586

YA0587

YA0588

CH3-

CH3~

CH3-

CH3-

СН3-

CH3-

CH3-





No.	R1	R2	R3	R4
YA0589	снз-	O <sub>2</sub> N-{}	н	СНЗ
YA0590	снз-	CN	н	СНЗ
YA0591	CH3-	NC	н.	CH3
YA0592	снз-	NC-(	Н	СНЗ
YA0593	снз-	CF₃	Н	СНЗ
YA0594	снз-	F₃C ⟨_)→₁	н	СНЗ
YA0595	снз-	F <sub>3</sub> C-{_}	н	СНЗ
YA0596	CH3	COOH	н	СНЗ
YA0597	СН3-	HOOC	н	СНЗ
YA0598	СН3	H00C-{_}-{	н	СНЗ
YA0599	CH3-	CO₂Me	н	снз
YA0600	CH3-	MeO₂C ⟨}-{	н	снз
YA0601	CH3	MeO <sub>2</sub> C-{}	н	СНЗ
YA0602	CH3-	CO₂Et	н	снз
YA0603	СН3-	EtO <sub>2</sub> C	н	снз
YA0604	CH3-	EtO <sub>2</sub> C-⟨}-{	Н	СНЗ
YA0605	снз-	SMe	. Н	снз
YA0606	СН3-	MeS	н	снз
YA0607	CH3-	MeS-{_}_	н	СНЗ
YA0608	CH3-	SO <sub>2</sub> Me	Н	СНЗ
YA0609	СН3-	MeO <sub>2</sub> S	н	СНЗ





No.	R1	R2_	R3	R4
YA0610	СН3-	MeO <sub>2</sub> S-{	Н	СНЗ
YA0611	СН3-	NH₂ ⟨¯⟩–₹	Н	СНЗ
YA0612	снз-	H <sub>2</sub> N	н	СНЗ
YA0613	снз-	H <sub>2</sub> N-{}-{	н	СНЗ
YA0614	CH3-	NMe₂	н	CH3
YA0615	снз-	Me <sub>2</sub> N	н	СН3
YA0616	снз-	Me <sub>2</sub> N-{	н	СНЗ
YA0617	снз~		Н	СН3
YA0618	снз-		н	СНЗ
YA0619	снз-		н	СНЗ
YA0620	снз-		н	СН3
YA0621	снз-		н	СН3
YA0622	СН3-		н	СНЗ
YA0623	СН3		H	СНЗ
YA0624	снз–		н	. СН3
YA0625	СН3-		Н	СНЗ
YA0626	СН3-	H3CN N-	Н	СНЗ
YA0627	СН3-	H³CN N-C	н	снз
YA0628	СН3-	H3CN_N-{}-{}	н	СНЗ
YA0629	СН3-	H <sub>3</sub> C CH <sub>3</sub>	Н	СНЗ
YA0630	CH3-	H³C-{\bigc}-	н	СНЗ

No.	RI	R2	R3	
YA0631		CH <sub>3</sub>	Н	CH3
YA0632	СН3-	CH₃ CH₃	н	СНЗ
YA0633	CH3-	H <sub>3</sub> C-\}-{	н	СНЗ
YA0634	снз-	H₃C H₃C	Н	СНЗ
YA0635	CH3-	F F	н	СНЗ
YA0636	CH3-	F-(S)-;	н	СНЗ
YA0637	СН3-	Ş F	н	СНЗ
YA0638	СН3-	€ F	Н	СНЗ
YA0639	СН3	F—	H	CH3
YA0640	СН3-	Ę <b>→</b> F	н	CH3
YA0641	СН3-	a_a ⇒	н	СНЗ
YA0642	СН3-	cı—(C)	H	СНЗ
YA0643	снз-	CI CI	, н	СНЗ
YA0644	CH3-	a G a	н	снз

No.	R1	R2	R3	R4
YA0645	СН3	d d d	н	снз
YA0646	СН3-	a √a	Н	СНЗ
YA0647	СН3-	H3CO_OCH3	Н	СН3
YA0648	СН3~	OCH <sub>3</sub> H₃CO-{_}}	н	СНЗ
YA0649	СН3-	OCH₃ → H₃CO	Н	снз
YA0650	СН3-	OCH <sub>3</sub> OCH <sub>3</sub>	н	СНЗ
YA0651	СН3-	H₃CO-⟨¯¯)}	Н	СНЗ

No.	R1	T		
110.	<del> ^-</del>	R2	R3	R4
YA0652	СН3~	H₃∞ H₃∞	н	СНЗ
YA0653	СН3-	F_OCH <sub>3</sub>	н	СНЗ
YA0654	CH3-	OCH <sub>3</sub>	н	СНЗ
YA0655	СН3-	OCH <sub>3</sub>	н	СНЗ
YA0656	СН3-	OCH3	н	СНЗ
YA0657	СН3-	OCH <sub>3</sub>	Н	СНЗ
YA0658	СН3-	OCH₃ F	н	. СНЗ
YA0659	СН3-	H₃CQ F—(□)—{	Н	СНЗ
YA0660	СН3-	H₃CO F	Н	СНЗ
YA0661	СН3-	H₃CO_F	н	СНЗ
YA0662	СН3-	H₃CO-⟨¯¯⟩ <del></del> }	н	СНЗ
YA0663	CH3-	H₃CO H₃CO	Н	СНЗ
YA0664	СН3	F, H₃CO-{_}}-}	н	СНЗ
YA0665	СН3-	CI_OCH₃	н	СНЗ

No.	R1	R2	R3	R4
YA0666	CH3-	OCH <sub>3</sub>	н	снз
YA0667	СН3-	OCH₃ G	н	снз
YA0668	СН3-	OCH <sub>3</sub> CI	H-	СНЗ
YA0669	снз-	G-{_}}-	Н	снз
YA0670	снз-	H₃CQ CD CD	н	снз
YA0671	снз-	H₃CO_CI →	н	СНЗ
YA0672	СН3-		н	СНЗ

No.	R1	R2	R3	R4
YA0673		H <sub>3</sub> CO	Н	CH3
YA0674	СН3-	H <sub>3</sub> CO-	Н	СНЗ
YA0675	СН3-	F_CH <sub>3</sub>	Н	СНЗ
YA0676	СН3-	CH <sub>3</sub>	н	СНЗ
YA0677	СН3-	CH <sub>3</sub> F	н	снз
YA0678	СН3-	CH <sub>3</sub>	н	СНЗ
YA0679	СН3-	H <sub>3</sub> C F—{}	н	СНЗ
YA0680	СН3-	H₃C F	н	СНЗ
YA0681	СН3	H₃C F	н	СН3
YA0682	CH3-	H₃C-⟨SH	н	СНЗ
YA0683	СН3-	H <sub>3</sub> C	Н	СНЗ
YA0684	СН3-	F_ H₃C-⟨¯¯);	н	СНЗ
YA0685	CH3-	Br_OCH <sub>3</sub>	Н	СНЗ
YA0686	CH3-	OCH <sub>3</sub>	н	снз

No.	R1	R2	R3	R4
YA0687	СН3-	OCH <sub>3</sub>	Н	снз
YA0688	снз-	OCH₃ Br	н	СНЗ
YA0689	СН3-	H₃CQ Br—⟨¯_)—;	Н.	снз
YA0690	СН3-	H₃CO Br	H	снз
YA0691	СН3-	H₃CO_Br	Н	снз
YA0692	СН3-	Br H₃CO-	н	CH3
YA0693	СН3-	Br ⊕} H₃co	Н	CH3

	F 64		1 5-	
No.	R1	R2	R3	R4
YA0694	СН3-	H³CO-⟨	н	СНЗ
YA0695	СН3-	H <sub>3</sub> CQ >	Н	СНЗ
YA0696	СН3-	OCH <sub>3</sub>	н	СНЗ
YA0697	СН3-	€ OCH3	н	СНЗ
YA0698	CH3-	H <sub>3</sub> CO_}	Н	СНЗ
YA0699	СН3-	H₃CO N-()	Н	СНЗ
YA0700	СН3-	CN OCH3	н	СНЗ
YA0701	СН3-	F F F	н	СНЗ
YA0702	СН3-	OCH₃ F-{_}} F	Н	СНЗ
· YA0703	СН3-	H₃CO-{_}} F	Н	СНЗ
YA0704	СН3-	OCH₃ F-⟨_}→} OCH₃	Н	СНЗ
YA0705	СН3-	OCH <sub>3</sub> OCH <sub>3</sub>	Н	СНЗ
YA0706	СН3-	a a—(a	Н	СНЗ
YA0707	снз-	CH-{SH3 CH3 CH3	Н	СН3



# PCT/JP2004/004320

No.	R1	R2	R3	R4
YA0708	СН3-	H₃co-{_}; a	н	снз
YA0709	СН3-	OCH3	Н	снз
YA0710	СН3-	OCH <sub>3</sub> H <sub>3</sub> CO-⟨∑→  OCH <sub>3</sub>	н	СН3
YA0711	СН3-	OCH <sub>3</sub>	н	CH3
YA0712	СН3-	H₃CO <del></del> <del></del>	Н	СНЗ
YA0713	СН3-	H₃CO-{_}-{_}-{_}-{_}-{_}-{_}-{_}-{_}-{_}-{_}	Н	СНЗ
YA0714	CH3-	OCH <sub>3</sub> }:	н	СНЗ

<u></u>	1 54		· · · · · · · · · · · · · · · · · · ·	
No.	R1	R2	R3	R4
YA0715	СН3-	H <sub>3</sub> CO	н	СНЗ
YA0716	СН3-	н₃со-⟨∑>-⟨∑\	Н	СНЗ
YA0717	СН3-	OCH <sub>3</sub>	н	СНЗ
YA0718	CH3-	H <sub>3</sub> CO	н	снз
YA0719	СН3-	H₃CO-<}_	н	снз
YA0720	СН3-	<b>₫-</b>	Н	снз
YA0721	СН3-	F	н	снз
YA0722	СН3-	F-()-()-!	н	СНЗ
YA0723	СН3-	5	н	снз
YA0724	CH3-		Н	СНЗ
YA0725	СН3-	F()-()	Н	СНЗ
YA0726	СН3-	<b>∅</b> - <b>√</b>	Н	СНЗ
YA0727	CH3-		н	СНЗ
YA0728	CH3-		н	СНЗ

No.	R1	R2	R3	R4
YA0729	СН3-		н	снз
YA0730	СН3-	CCC,	н	снз
YA0731	CH3-	СН3-	н	Q
YA0732	CH3-	CH3CH2-	н	Q.
YA0733	СН3-	<b>^</b> \	Н	Q
YA0734	СН3-	\\\\\	н	Q
YA0735	СН3-	<b>\\\\</b>	н	Q

N-	I D1	T		
No.	R1	R2	R3	R4
YA0736	снз-		н	Q
YA0737	снз-	$\gamma$	Н	Q
YA0738	снз-	Y	Н	Q
YA0739	СН3-	~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Q
YA0740	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Q
YA0741	СН3-	X.	н	Q
YA0742	СН3-	7	Н	Qu
YA0743	снз-	<b>\\\\</b>	н	Qu
YA0744	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Q
YA0745	СН3-	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	. н	Qu
YA0746	СН3-	<b>\\\\</b>	н	Q
YA0747	СН3-	<b>\\\\\</b>	н	Q
YA0748	CH3-		Н	
YA0749	СН3-		н	
YA0750	СН3-		н	Qu
YA0751	СН3-		Н	
YA0752	снз-	$\triangleright$	Н	Qu
YA0753	CH3-	$\Diamond$ -I	Н	Qu
YA0754	СН3-	$\bigcirc$	Н	Qu
YA0755	СН3-	$\supset$	н	Qu
YA0756	снз-	$\bigcirc$ $\dashv$	н	

No.	R1	R2	T D2	
YA0757	CH3-		R3 H	R4
YA0758	СН3-		Н	Q
YA0759	CH3-	<u></u> m.{	Н	Qu
YA0760	СН3-	F	н	Q
'YA0761	снз-	<u></u>	н	Q
YA0762	снз-		н	Q
YA0763	снз-	F-(-)	н	Qi
YA0764	снз-	F-()(	Н	
YA0765	CH3-	CI C	н	
· YA0766	СН3-	CI	Н	
YA0767	СН3-	CH	н	
YA0768	снз-	c⊢ <b>(</b> )→	Н	
YA0769	СН3-	CI—(	Н	
YA0770	СН3-	Br ∤	Н	
YA0771	CH3-		Н	
YA0772	CH3-	Br-{_}}	H	
YA0773	CH3-	Br- <b>(</b> ){	. н	Q
YA0774	CH3-	Br{	н	
YA0775	CH3-		н	Q
YA0776	СН3-		н	Qu
YA0777	снз-	H	н	Q

		<u> </u>	•
No.	R1	R2	R3 R4
YA077	8 снз-	<u>{_</u> } <del>-</del>	н
YA0779	СН3-	H <sub>3</sub> C	н
YA0780	снз-	H <sub>3</sub> C-{}	H
YA0781	CH3-	C <sub>2</sub> H <sub>5</sub> -{}	н
YA0782	СН3-	n-C <sub>3</sub> H <sub>7</sub> -{}{	н
YA0783	СН3-	л-С <sub>4</sub> Н <sub>9</sub> -{}-{	н
YA0784	CH3-	OH OH	H
YA0785	СН3-	HO	н
YA0786	CH3-	HO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar	н
YA0787	СН3-	OCH₃	H
YA0788	CH3-	H³co	H
YA0789	СН3-	H <sub>3</sub> CO-{}-{	н
YA0790	СН3-	H <sub>3</sub> CO-{_}{	н
YA0791	СН3-	H <sub>3</sub> CO-{\bigs\n.{}	н
YA0792	CH3-	OC <sub>2</sub> H <sub>5</sub>	Н
YA0793	СН3-	C <sub>2</sub> H <sub>5</sub> O	н
YA0794	CH3-	C <sub>2</sub> H <sub>5</sub> O-{_}-	. н 🔍
YA0795	СН3-	n-C₃H <sub>7</sub> O-⟨}~-{	H
YA0796	СН3-	n-C₄H₀O-⟨_ <mark>⟩</mark> {	н
YA0797	СН3-	NO <sub>2</sub>	H
YA0798 ·	СН3-	O <sub>2</sub> N	н
			·

No.	R1	R2		
YA0799		0.11	R3 H	R4
YA0800	снз-	CN	н	Q
YA0801	CH3-	NC	Н	Q
YA0802	снз-	NC-	н	Qu
YA0803	снз-	CF <sub>3</sub>	н	Qu
YA0804	CH3-	F <sub>3</sub> C	н	Q
YA0805	СН3-	F <sub>3</sub> C-{_}-{	Н	Q
YA0806	CH3-	COOH	Н	Q
YA0807	СН3-	HOOC	н	Q
YA0808	CH3-	H00C-{\rightarrow}-1	н	
YA0809	СН3-	CO <sub>2</sub> Me	н	Q
YA0810	СН3-	MeO <sub>2</sub> C	Н	Q
YA0811	снз-	MeO <sub>2</sub> C-{}	н	
YA0812	CH3-	CO₂Et	н	Q
YA0813	СН3-	EtO <sub>2</sub> C	н	Q
YA0814	СН3-	EtO <sub>2</sub> C-{}_{_}	н	Q
YA0815	СН3-	SMe	Н -	Q
YA0816	CH3-	MeS \_\_	Н	
YA0817	CH3-	MeS-{_}	н (	1
YA0818	СН3-	SO <sub>2</sub> Me	н	2
YA0819	СН3-	MeO <sub>2</sub> S	н (	



No.	- 6.		
NO.	R1	R2	R3 R4
YA082	0 снз-		н
YA0821	СН3-		н
YA0822	2 СН3-	H <sub>2</sub> N	н
YA0823	СН3-		н
YA0824	СН3-		н
YA0825	СН3-	Me <sub>2</sub> N	н
YA0826	СН3-	Me <sub>2</sub> N-	H Q
YA0827	снз-		н
YA0828	снз-		н
YA0829	CH3-	Cn-€]-1	н
YA0830	СН3-	○+	н
YA0831	CH3-		н
YA0832	CH3-	(N-()-1	н
YA0833	CH3-	0_N <del>-</del> (_)	н
YA0834	CH3-		н
YA0835	снз-	<b>○</b> N-()-}	н 🔍
YA0836	СН3-	H³CN_N-⟨	. н 🔍
YA0837	СН3-	H3CN N-	H
YA0838	снз-	H3CN N-{}	н
YA0839	СН3-	H <sub>3</sub> C_CH <sub>3</sub>	Н
YA0840	CH3-	CH₃ H₃C-⟨¯}→¦	н

YA0851

YA0852

YA0853

YA0854

YA0855

СН3-

СН3-

СН3--

CH3-

СН3-

1/085408				PCT/JP2
No.	R1	R2	R3	R4
YA0841	снз-	CH₃ CH₃ H₃C	н	Qu
YA0842	снз-	CH₃ CH₃	Н	Qu
YA0843	СН3-	H <sub>3</sub> C-(	н	Qu
YA0844	СН3-	H₃C H₃C	н	Qu
YA0845	CH3-	F F	н	
YA0846	СН3	F-⟨¯¯⟩-;	н	Qu
YA0847	СН3-	ŞŠ-; F	н	
YA0848	CH3-	F F	н	
YA0849	СН3-	F————;	Н	Qu
YA0850	снз-	F	н	Qu

185

Н

Н

н

н

Н

No.	R1			
140.	<del> </del>	R2	R3	R4
YA0856	CH3-	G G	н	Q
YA0857	снз-	H₃CO_OCH₃	Н	Qu
YA0858	СН3-	осн <sub>3</sub> н₃со-⟨\$\;	Н	Qu
YA0859	CH3-	OCH <sub>3</sub> H <sub>3</sub> CO	н	
YA0860	СН3-	OCH <sub>3</sub> .	Н	
YA0861	CH3-	H₃CO-⟨¯_}-}	н	Q

No.	L RI	R2	T 50	
	<u> </u>	H <sub>3</sub> CQ	R3	R4
YA0862	СН3-	H <sub>3</sub> CO	н	
YA0863	СН3-	F_OCH <sub>3</sub>	н	Qu
YA0864	СН3-	OCH <sub>3</sub>	. н	Q
YA0865	СН3~	OCH <sub>3</sub>	Н	Qu
YA0866	СН3-	OCH <sub>3</sub> F—∑iii√	H	Qu
YA0867	СН3-	OCH₃ ↓ F	н	Q
YA0868	СН3-	OCH <sub>3</sub> F	н	Q
YA0869	СН3-	H₃CO F—√—}	Н	Qu
YA0870	СН3-	H₃CO F	Н	
YA0871	CH3-	H₃CO_F	Н	
YA0872	CH3-	H₃CO-{	Н	
YA0873	СН3-	H³CO F	Н	
YA0874	СН3-	F. H₃CO-⟨□}∤	H	Q
YA0875	СН3	CI_OCH <sub>3</sub>	н	Qu
YA0876	СН3-	CI—{_}→} OCH³	Н	Qi



### PCT/JP2004/004320

No.	R1	R2	R3	R4
YA0877	снз-	OCH₃ G	Н	Qu
YA0878	снз-	ocH <sub>3</sub> ⇔	Н	Q
YA0879	CH3-	H₃CO CI—()—;	Н	Q
YA0880	СН3-	H₃cq	н	Q
YA0881	СН3-	H₃CO_CI	н	Q
YA0882	CH3-	CI H₃CO-{}}	Н	Q

No.	R1		T	
7.0.	- NI	R2	R3	R4
YA0883	CH3-	H <sub>3</sub> CO	н	
YA0884	СН3-	CI, H₃CO-{\( \)}—{	Н	Q
YA0885	СН3-	F_CH <sub>3</sub>	Н	Qu
YA0886	СН3-	CH₃ F—{\bar{\bar{\bar{\bar{\bar{\bar{\bar	Н	Q
YA0887	СН3-	CH <sub>3</sub> F	Н.	Q
YA0888	СН3-	CH₃	н	Q
YA0889	СН3	H₃C F-{}	Н	
YA0890	CH3-	H₃C }_ F	Н	
YA0891	СН3-	H₃C_F	н	Qu
YA0892	снз-	H₃C-⟨S}-	Н	Qu
YA0893	снз-	F H₃C	Н	Qu
YA0894	снз-	F_ H₃C-⟨¯}→	Н	Q
YA0895	СН3-	Br_OCH <sub>3</sub>	н	Qu
YA0896	СН3-	OCH₃ Br—⟨□}—{	н	Qu
YA0897	СН3-	OCH₃ Br	н	Q

No.	R1	R2	T	T
YA0898	CH3-	OCH <sub>3</sub> Br	R3 H	R4
YA0899	СН3	H₃CQ Br—()—;	Н	Q
YA0900	СН3-	H₃CQ Br	Н	Q
YA0901	СН3-	H₃CQ_Br	н	Q
YA0902	СН3	Br H₃CO-⟨¯¯ <mark>&gt;</mark> }	н	Q
YA0903	CH3-	H <sub>3</sub> CO	Н	Qu

No.	Ri	R2	1 00	<del></del>
YA0904		Br H <sub>3</sub> CO-	R3 H	R4
YA0905	CH3-	H <sub>3</sub> CO >	Н	Qi
YA0906	СН3-	CN-CDH3	Н	Que
YA0907	снз-	CN-C}-OCH₃	н	Q
YA0908	СН3	H <sub>3</sub> CO	Н	Q
YA0909	CH3~	H³CO H³CO	н	Q.
YA0910	СН3-	CN CH₃	Н	Q
YA0911	CH3-	F-()-;	н	Q
YA0912	СН3-	OCH <sub>3</sub> F—{_}} F	Н	
YA0913	CH3-	H₃CO-{-\$-{ F	Ħ	
YA0914	CH3-	OCH₃ F—C OCH₃	Н	Qu
YA0915	СН3-	OCH <sub>3</sub> H₃CO-⟨_} OCH <sub>3</sub>	н	Qu
YA0916	снз-	a — (a a	Н	Q <sub>1</sub>
YA0917	СН3-	OCH <sub>3</sub> CI-{_}}-{ CI	н	Qi
YA0918	СН3-	H³co-{∑} CI	н	Q

188

CLIDIADAMAN VALLERANDO CAR





## PCT/JP2004/004320

No.	R1	R2	R3	R4
YA0919	СН3-	OCH <sup>2</sup> OCH <sup>3</sup>	Н	
YA0920	СН3-	OCH3 OCH3	Н	Q
YA0921	CH3-	OCH₃	н	Q
YA0922	CH3-	H₃CO ————	н	Q
YA0923	СН3	H3CO-{\}-{\}-{\}-{\}	Н	Q
YA0924	СН3-	OCH <sub>3</sub> \t	н	

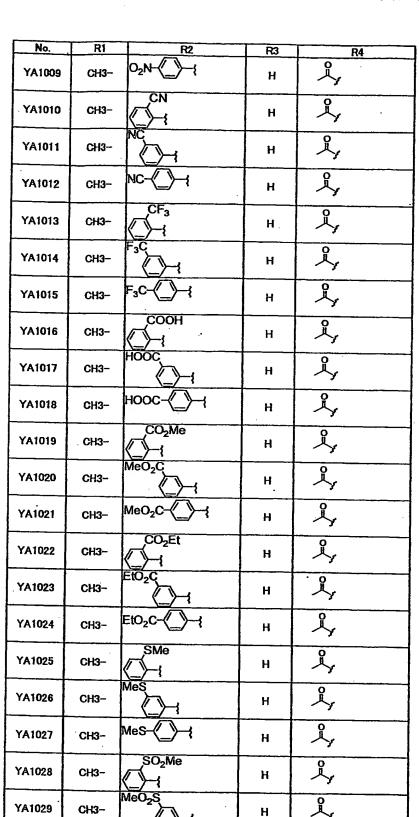
No.	R1	7	T	
140.	<del>  ^'</del>	R2	R3	R4
YA0925	СН3-	H <sub>3</sub> CO	н	
YA0926	СН3-	H3CO-{\( \)\( \)_\tag{\chi}	н	Q
YA0927	CH3-	OCH <sub>3</sub>	н	Q
YA0928	СН3-	H <sub>3</sub> CQ	н	Q
YA0929	СН3-	н₃со-⟨}	Н	Q
YA0930	СН3-	₫-O-;	н	Qu
YA0931	СН3-	F	Н	Q
YA0932	СН3-	F-{\}-{\}-{\}-	Н	Qu
YA0933	СН3-	ď-ď	н	
YA0934	CH3-		н	Q
YA0935	СН3-	F-()-()	Н	
YA0936	CH3-		Н	Q
YA0937	СН3-		н	Q
YA0938	СН3-	F-()-()	Н	Q
YA0939	СН3-		Н	Q

No.	T D1	γ		
140.	R1	R2	R3	R4
YA0940	СН3-	CC,	Н	Q
YA0941	CH3-	CH3	Н	l,
YA0942 <sub>.</sub>	СН3-	СНЗСН2-	н	L,
YA0943	СН3-	<b>^</b> ∖\	Н	<u>گ</u>
YA0944	СН3	Y	H.	Ŷ,
YA0945	СН3	<b>√</b> \	н	L,

No.	R1	R2	1 200	
YA0946			R3 H	R4
YA0947	снз-	7	Н	Î,
YA0948	снз-	丫	Н	l,
YA0949	снз-	~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Å,
YA0950	СН3-	Y~	н	الم الم
YA0951	СН3-	X	н	Î,
YA0952	CH3-	7	Н	Å,
YA0953	СН3-	~~~``	н	Ŷ,
YA0954	CH3-		Н	Ŷ,
YA0955	CH3-	~~~\	н	Ŷ,
YA0956	CH3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	بُ
YA0957	CH3-	~~~``	Н	Ŝ,
YA0958	СН3-		н	بُ
YA0959	CH3-		н	Å,
YA0960	CH3-		н	بُ
YA0961	СН3		н	ئى
YA0962	СН3-	<b>▷</b> →	Н	<u></u>
YA0963	CH3-	$\Diamond$ -I	н	گي
YA0964	снз-	$\bigcirc$	Н	گہ
YA0965	CH3-	$\bigcirc$ $\vdash$	н	Ŷ,
YA0966	CH3-	$\bigcirc$ H	н	2,

No.	R1	R2	R3	R4
YA0967	CH3-		Н	Ŷ,
YA0968	СН3-		н	<u></u>
YA0969	СН3-	<b>⊘</b> {	н	Î,
YA0970	СН3-	Ç,	Н	Ŷ,
YA0971	CH3-		н	Ŷ,
YA0972	СН3-	F()-1	н	Ŷ,
YA0973	СН3-		н	Ŷ,
YA0974	СН3-		н	Ŷ,
YA0975	СН3-		н	Ŷ,
YA0976	СН3-	CI ————————————————————————————————————	н	Ŷ,
YA0977	CH3-	c⊢ <b>(_</b> }-{	н	,
YA0978	CH3-	C <b>├</b> ───	н	<u></u>
YA0979	CH3	C⊢-(_)n·∤	н	<u></u>
YA0980	CH3-	Br	Н	
YA0981	CH3-	Br.	н	
YA0982	CH3-	Br-{}-{	н	<u></u>
YA0983	CH3-	Br-{}-{	н	2,
YA0984	CH3-	Br—(	н	L <sub>y</sub>
YA0985	снз-		н	<u>}</u> ,
YA0986	СН3-		н	Å,
YA0987	снз-		Н	2,

No.	R1	R2	R3	R4
NO.	- 111	CH₃	-10	
YA0988	СН3-		н	l,
YA0989	СН3-	H₃C ———	Н	Ļ,
YA0990	СН3-	H3C- <b>(</b> ){	Н	Ļ,
YA0991	CH3-	C <sub>2</sub> H <sub>5</sub> -{}-{	н	Ļ,
YA0992	снз-	n-C <sub>3</sub> H <sub>7</sub> -{\rightarrow}	Н	Å,
YA0993	CH3	n-C <sub>4</sub> H <sub>9</sub> {}{	Н	Ŷ,
YA0994	CH3-	OH OH	н	ڳ
YA0995	CH3-	HO HO	н	Ŷ,
YA0996	снз-	HO-{}-}	н	Ļ,
YA0997	СН3-	OCH <sub>3</sub>	н	Ĵ,
YA0998	CH3-	H <sub>3</sub> CO	н	Å,
YA0999	CH3-	H <sub>3</sub> CO-{_}-{	н	Å,
YA1000	CH3-	H <sub>3</sub> CO-{}-{	н	<u>ئ</u>
YA1001	СН3-	H <sub>3</sub> CO-{\bigs\nu}!"\f	н	<u></u>
YA1002	СН3-	OC <sub>2</sub> H <sub>5</sub>	н	<u></u>
YA1003	снз-	C <sub>2</sub> H <sub>5</sub> O	н	以
YA1004	СН3-	C <sub>2</sub> H <sub>5</sub> O-{}	н	l,
YA1005	СН3-	n-C <sub>3</sub> H <sub>7</sub> O-{}{1	н	Î,
YA1006	СН3-	n-C₄H <sub>9</sub> O- <b>⟨</b> }-{	н	l,
YA1007	СН3-	NO <sub>2</sub>	н	گہ
YA1008	СН3-	O <sub>2</sub> N	н	<u>گ</u>



No.	. R1	R2	R3	R4
YA1030	СН3-	MeO <sub>2</sub> S-{	Н	Å,
YA1031	СН3-	NH <sub>2</sub>	н	l,
YA1032	СН3-	H2N	Н	Î,
YA1033	СН3-	H <sub>2</sub> N-{}-{	н	<u>گ</u>
YA1034	снз-	NMe <sub>2</sub>	Н	گ <sub>ا</sub>
YA1035	СН3-	Me <sub>2</sub> N	н	Ŷ,
YA1036	CH3-	Me <sub>2</sub> N-	н	بُ
YA1037	снз-		н	Ŷ,
YA1038	СН3-		н	<u></u>
YA1039	СН3-		н	Å,
YA1040	СН3-		н	Å,
YA1041	СН3-		Н	3
YA1042	СН3-		.Н	گہ
YA1043	CH3-		н	l,
YA1044	CH3-		н	Ž,
YA1045	СН3-	< <u></u> \-\_}-	. н	l,
YA1046	CH3-	H3CN_N-{\}	. н	Ĵ,
YA1047	CH3-	H3CN_N-{}	Н	Î,
YA1048	0.10	H <sub>9</sub> CN_N-{}-{	н	گې
YA1049	CH3-	H₃C_CH₃ ⟨∑	н	Ŷ,
YA1050	снз-	CH₃ H₃C-{{}}-{	Н	گ,

	1 5/			
No.	R1	R2	R3	R4
YA1051	СН3-	CH₃	н	l,
YA1052	СН3-	CH <sub>3</sub>	Н	Î,
YA1053	СН3-	H <sub>3</sub> C H <sub>3</sub> C-\}	н	Ŷ,
YA1054	СН3-	H₃C H₃C	Н	Ŷ,
YA1055	СН3-	F_F	Н	Å,
YA1056	СН3-	F F	Н	Ŷ,
YA1057	СН3-	Ş F F	Н	l,
YA1058	CH3~	F F	н	Ŷ,
YA1059	СН3-	F	Н	Ŷ,
YA1060	СН3-	F.	Н	l,
YA1061	СН3-	a_a	н.	l,
YA1062	снз-	a—⟨	Н	Ŷ,
YA1063	. СН3-	a a	Н	l,
YA1064	снз-	a √a	Н	L,

No.	R1	R2	R3	R4
YA1065	СН3	a, a,	Н	l,
YA1066	снз-	a a	н	Å,
YA1067	CH3-	H₃CO_OCH₃	Н	l,
YA1068	СН3-	OCH <sub>3</sub>	Н	Ŷ,
YA1069	СН3-	H³CO H³CO	н	l,
YA1070	СН3-	OCH3	н	Ŷ,
YA1071	СН3-	H₃CO H₃CO-⟨¯}}	н	l,

No.	R1	T		
YA1072	CH3-	H <sub>3</sub> CO	R3 H	R4
YA1073	снз-	F_OCH <sub>3</sub>	Н	گړ
YA1074	СН3-	OCH <sub>3</sub>	н	l,
YA1075	СН3	OCH <sub>3</sub>	Н	l,
YA1076	снз-	OCH <sub>3</sub>	н	l,
YA1077	СН3-	OCH₃ F	н	Ŷ,
YA1078	СН3-	OCH₃ F	H	Ŷ,
YA1079	снз-	H₃CO F—()—{	Н	Ŷ,
YA1080	CH3-	H₃CO F	Н .	Ŷ,
YA1081	СН3-	H <sub>3</sub> CO_F	Н	Ŷ,
YA1082	снз-	F H₃CO-{\bigc\}}	Н	l,
YA1083	СН3-	F H₃∞	н	Ļ,
YA1084	СН3-	H₃CO-{_}}-{	н	Ž,
YA1085	CH3-	CI_OCH₃	H -	Ļ

No.	R1	R2	R3	R4
YA1086	СН3-	CH-€_}	Н	Å,
YA1087	СН3-	OCH <sub>3</sub>	н	Å,
YA1088	СН3-	CI OCH3	н	Ŷ,
YA1089	СН3-	H₃CQ CI—⟨∑}—;	Н	Å,
YA1090	СН3	H₃Q C	Н	Å,
YA1091	CH3	H <sub>3</sub> CO_CI	н	l,
YA1092	СН3	CI H₃CO-⟨¯_}—;	Н	Ŷ,

No.	RI	D2	D2 3	D4
140.	- KI	R2	R3	R4
YA1093	CH3-	H <sub>3</sub> CO	н	
YA1094	CH3-	CI H₃CO-{}}	н	Î,
YA1095	снз-	F_CH <sub>3</sub>	н	l,
YA1096	CH3-	CH₃ F—⟨\$\rightarrow\$\rightarrow\$	Н	Ŷ,
YA1097	CH3-	CH₃ F	Н	Ŷ,
YA1098	CH3-	CH₃ ← F	Н	Ŷ,
YA1099	CH3-	H <sub>3</sub> C F-\(\begin{array}{c}\)-\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	Ŷ,
YA1100	CH3-	H₃C → F	н	l,
YA1101	CH3-	H₃C_F	н	Ŷ,
YA1102	CH3-	F H₃C-⟨¯_}→	н	Ŷ,
YA1103	CH3-	F H₃C	Н	l,
YA1104	CH3-	F H₃C-{\}{	н	l,
YA1105	CH3-	Br_OCH <sub>3</sub>	н	Å,
YA1106	СН3-	OCH₃ Br—{}	н	l,

r		T		
No.	R1	R2	R3	R4
YA1107	снз-	OCH <sub>3</sub> Br	н	l,
YA1108	СН3	OCH <sub>3</sub> Br	Н	l,
YA1109	CH3-	H <sub>3</sub> CO Br	н	l,
YA1110	СН3-	H₃CO Br	Н	Ŷ,
YA1111	CH3	H₃CO_Br	н	Ŷ,
YA1112	CH3-	Br H₃CO-{\bar{\bar{\bar{\bar{\bar{\bar{\bar	н.	Ŷ,
YA1113	СН3-	Br H₃CO	Н	\$



No.	l D1			
140.	R1	R2	R3	R4
YA1114	СН3-	H³CO-{_}	н	l,
YA1115	СН3-		Н	Ŷ,
YA1116	СН3-	OCH <sub>3</sub>	Н	, L,
YA1117	СН3-	CN-C}ocH₃	н	l,
YA1118	CH3-	H <sub>3</sub> CO_}-N	н	l,
YA1119	СН3-	H₃CO N-√□-	н	l,
YA1120	СН3-	CN OCH³	н	Å,
YA1121	СН3-	F-{-}  F	н	Å,
YA1122	СН3-	OCH₃ F—{_}} F	Н	l,
YA1123	СН3	H₃CO-{\}_F F	н	Ŷ,
YA1124	СН3	OCH <sub>3</sub> F-(_) OCH <sub>3</sub>	н	l,
YA1125	СН3-	OCH3	н	l,
YA1126	СН3-	a—{∑i a	н	L,
YA1127	СН3-	ocH₃ cr Cr Cr	Н	l,

No.	R1	R2	R3	R4
YA1128	СН3-	H₃CO-{_}}; CI	Н	Ŷ,
YA1129	СН3-	OCH <sup>3</sup> CH-{_}} OCH <sup>3</sup>	Н	Ŷ,
YA1130	СН3-	OCH <sub>3</sub> H <sub>3</sub> CO-⟨_∑-{ OCH <sub>3</sub>	Н	Ŷ,
YA1131	СН3~	OCH <sub>3</sub>	Н	l,
YA1132	СН3-	H₃CO	Н	2,
YA1133	СН3-	н₃со-⟨_}-⟨_}-∤	Н	Ŷ,
YA1134	СН3-	OCH37	Н	Å,

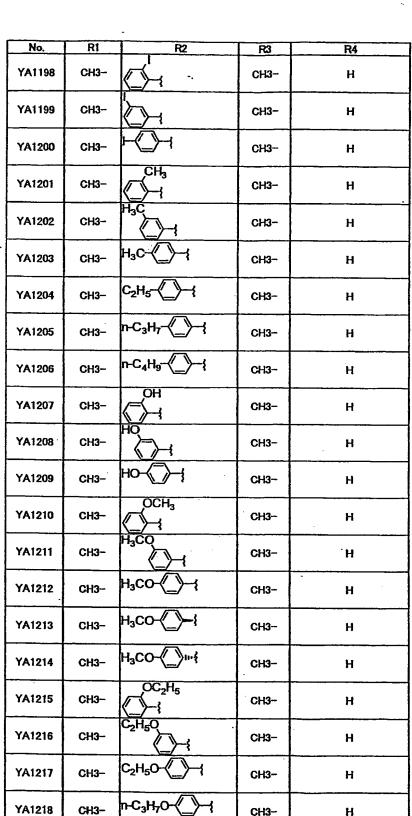
No.	Rt	T	1	•
YA1135	CH3-	H <sub>3</sub> CO ,	R3 H	R4
YA1136	CH3-	н₃со-⟨∑-⟨_∑\	Н	l,
YA1137	СН3-	OCH <sub>3</sub>	н	L,
YA1138	CH3-	H <sub>3</sub> CO	Н	l,
YA1139	СН3-	H <sub>3</sub> CO-{}	н	Ŷ,
YA1140	СН3~	<b>₽</b>	Н	Ŷ,
YA1141	СН3-	F	Н	l,
YA1142	СН3-	F-()-()-{	н	l,
YA1143	СН3		н	l,
YA1144	СН3-		Н	.L,
YA1145	СН3-	F-(-)-(-)	н	l,
YA1146	СН3-	F,	Н	l,
YA1147	СН3-		Н	Ŷ,
YA1148	СН3	F-Q-Q	н	Š,

No.	R1	R2	R3	R4
YA1149	СН3-	Q	н	l,
YA1150	снз-	CC,	н	Å,
YA1151	CH3-	CT)	Н	Å,
YA1152	СН3-	O's	н	Ŷ,
YA1153	СН3-	Ğ.	н	l,
YA1154	СН3-	снз-	CH3-	Н
YA1155	СН3-	CH3CH2-	СН3-	Н

No.	R1	.D2		
		R2	R3	R4
YA115	6 CH3-		CH3-	н
YA115	7 СН3-	Y	СН3-	, н
YA115	6 CH3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	снз–	Н
YA115	9 снз-		снз-	Н
YA1160	О СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	СН3-	н
YA1161	СН3	丫	СН3-	н
YA1162	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	СН3-	Н
YA1163	снз-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	СН3-	Н
YA1164	снз–	X.	СН3-	н
YA1165	снз-	7	СН3-	н
YA1166	CH3~	<b>\\\\\</b>	СН3-	н
YA1167	снз-	L~	СН3-	н
YA1168	CH3-	<b>^</b>	снз-	Н
YA1169	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	СН3-	н
YA1170	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	CH3-	Н
YA1171	СН3~		CH3-	н
YA1172	снз-	Q	СН3-	н
YA1173	снз-		СН3-	Н
YA1174	CH3-		СН3-	Н
YA1175	СН3-	$\triangleright$	CH3-	Н
YA1176	CH3-	$\Diamond$ -1	CH3-	н

# PCT/JP2004/004320

No	R1	R2	R3	R4
YA1177	СН3-	$\bigcirc$	CH3-	н
YA1178	снз-	$\bigcirc$ $\dashv$	CH3-	н
YA1179	СН3-	$\bigcirc$ $\dashv$	СН3-	н
YA1180	СН3-		CH3-	н
YA1181	СН3-		снз-	Н
YA1182	CH3-	<b></b>	СН3-	Н
YA1183	СН3-	F	CH3-	н
YA1184	СН3		CH3-	н
YA1185	СН3-	F-()-1	СН3-	Н
YA1186	снз-		снз–	Н
YA1187	СН3⊸	F—C)in {	СН3-	Н
YA1188	CH3	CI C)	СН3-	. н
YA1189	CH3-	CI ————————————————————————————————————	СН3-	н
YA1190	СН3-	c⊢ <b>(_</b> }- <sub>1</sub>	CH3-	н
YA1191	CH3-	c⊢∕_}-(	снз-	. н
YA1192	СН3-	CH	СН3	Н
YA1193	CH3-	Br	CH3-	Н
YA1194	СН3-	Br.	СН3-	Н
YA1195	СН3-	Br-{_}	CH3-	Н
YA1196	снз-	Br—{}	снз-	Н
YA1197	CH3-	Br—(	CH3-	н



CH3-

Н



No.	D1	T		
INO.	. R1	R2	R3	R4
YA1219	СН3-	n-C <sub>4</sub> H <sub>9</sub> O-(){	снз-	Н
YA1220	снз-	NO <sub>2</sub>	СН3-	Н
YA1221	CH3-	O <sub>2</sub> N	CH3-	н
YA1222	СН3-	O <sub>2</sub> N-()-(	CH3-	Н
YA1223	СН3-	CN	СН3-	Н
YA1224	СН3-	NC	CH3-	н
YA1225	CH3-	NC-{}	СН3-	Н
YA1226	СН3-	NH <sub>2</sub>	СН3-	н
YA1227	CH3-	H <sub>2</sub> N	СН3-	н
YA1228	СН3-	H <sub>2</sub> N-(T)(	снз-	н
YA1229	CH3-	NMe <sub>2</sub>	СН3-	Н
YA1230	СН3-	Me <sub>2</sub> N	СН3-	Н
YA1231	СН3-	Me <sub>2</sub> N-()-(	СН3-	Н
YA1232	СН3-	Cr-(2)	CH3-	н
YA1233	CH3-	O+Q	CH3	Н
YA1234	CH3-	(h-()-1	СН3-	Н
YA1235	CH3-		СН3-	н
YA1236	СН3-		снз-	Н
YA1237	СН3-		СН3-	н
YA1238	СН3-		СН3-	н
YA1239	СН3-		CH3-	н

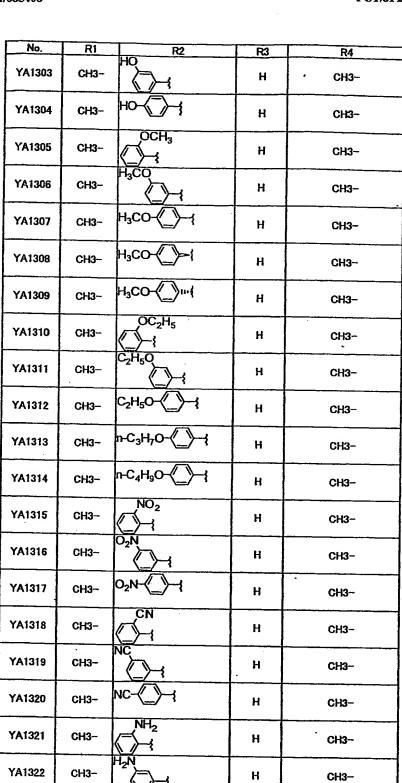
No.	R1	R2	R3	7.
YA1240		©N-{}-1	снз-	R4 H
YA1241	снз-	H3CN_N-	СН3-	Н
YA1242	CH3-	H3CN N-	CH3-	Н
YA1243	снз-	H3CN_N-{}-{	CH3-	н
YA1244	снз-	OCH <sub>3</sub>	CH3-	н
YA1245	снз-		СН3-	н
YA1246	снз-	OCH <sub>3</sub>	СН3-	н
YA1247	СН3-		CH3-	н
YA1248	CH3-	CCC,	CH3-	Н
YA1249	СН3-	снз	н	СН3-
YA1250	СН3-	СНЗСН2-	н	СН3-
YA1251	CH3-	<b>^</b> \	н	СН3-
YA1252	СН3-	Y	н	СН3
YA1253	CH3-	<b>\\\\</b>	н	СН3-
YA1254	CH3-	<u></u>	н	снз-
YA1255	CH3-	$ \uparrow $	н	CH3-
YA1256	CH3	丫	. н	СН3-
YA1257	СН3-	~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3-
YA1258	CH3-	~~	н	СН3-
YA1259	CH3-	r.k	н	СН3-
YA1260	CH3-	个	н	СН3-





No.	RI	Do	- D2	
	<del> '``</del> -	R2	R3	R4
YA1261	СН3-	~~~	Н	CH3-
YA1262	CH3-	L	Н	CH3-
YA1263	СН3-	~~~``\	٠н	СН3-
YA1264	снз-	~~~~	Н	СН3-
YA1265	СН3-	~~~``	Н	снз-
YA1266	снз–	L~~~	Н	СН3-
YA1267	снз-		н	СН3-
YA1268	СН3-		н	СН3-
YA1269	снз–		Н	СН3-
YA1270	CH3-	<b>→</b>	Н	СН3-
YA1271	снз-	$\Diamond$ -i	н	CH3-
YA1272	CH3-	$\bigcirc \dashv$	Н	CH3-
YA1273	CH3-		н	СН3-
YA1274	СН3-	$\bigcirc \dashv$	н	CH3-
YA1275	СН3-		. н	СН3-
YA1276	СН3-		н	СН3-
YA1277	СН3-	<u></u>	Н	снз-
YA1278	снз-	<u></u>	Н	СН3-
YA1279	снз-		н	CH3-
YA1280	снз-	F-()-1	Н	снз-
YA1281	CH3-	F-()-1	н	СН3-

No.	R1	R2		· · · · · · · · · · · · · · · · · · ·
<del>- , ,,,,-</del>	·   · · · ·		R3	R4
YA1282	CH3-	F-()11-4	н	снз-
YA1283	CH3-	CI C	н	СН3
YA1284	СН3-	CI	Н	СН3-
YA1285	снз-	CH	Н	снэ-
YA1286	снз-	CH	н	снз
YA1287	. СН3-	CH	н	CH3-
YA1288	снз-	Br △	н	снз-
YA1289	снз-	Br.	Н	снз-
YA1290	CH3-	Br-{}-{	Н	снз-
YA1291	CH3	Br-{\}-{	н	снз-
YA1292	снз-	Br—(	н	снз-
YA1293	CH3-		н	снз-
YA1294	СН3~		н	снз
YA1295	СН3~		н	снз-
YA1296	СН3-	CH <sub>3</sub>	н	СН3-
YA1297	СН3	H <sub>3</sub> C	Н	CH3-
YA1298	снз-	H₃C- <b>⟨</b>	. н	СН3-
YA1299	снз-	C <sub>2</sub> H <sub>5</sub> -{}-{	Н	CH3-
YA1300	СН3-	n-C₃H <sub>7</sub> {_}}{	н	CH3-
YA1301	СН3-	n-C <sub>4</sub> H <sub>9</sub> -{	н	СН3-
YA1302	снз-	OH ⟨□⟩	н	CH3-



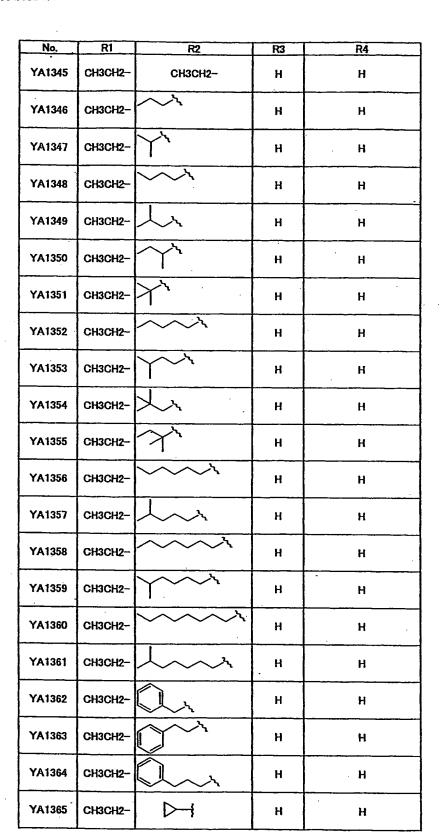
Н

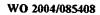
CH3-

YA1323

CH3-

No.	. R1	50		
140.		NMe <sub>2</sub>	R3	R4
YA1324	CH3-		н	СН3-
YA1325	снз-	Me <sub>2</sub> N	н	CH3-
YA1326	снз–	Me <sub>2</sub> N-(	Н	CH3-
YA1327	CH3-	CN-S	Н	снз-
YA1328	снз-	CHQ	н	снз-
YA1329	СН3-	CH-C)-1	Н	снз-
YA1330	снз-		Н	СН3-
YA1331	снз-		н	СН3-
YA1332	снз-		Н	СН3-
YA1333	СН3-		Н	CH3-
YA1334	СН3-		Н	снз-
YA1335	снз-		Н	снз-
YA1336	снз-	H3CN N-	н	CH3-
YA1337	CH3-	H³CN N-⟨\}	н	СН3
YA1338	СН3-	H³CN_N-{}	Н	. СН3-
YA1339	СН3-	F-€D-+	н	снз-
YA1340	CH3-	OCH <sub>3</sub> F-()-(	н	СН3-
YA1341	СН3-	OCH <sub>3</sub>	н	CH3-
YA1342	СН3-		н	СН3-
YA1343	CH3-		н	СН3-
YA1344	СНЗСН2-	снз-	н	Н







No.	RI	R2	1 m	T
YA1366		$\Diamond$	R3 H	R4
			"	Н
YA1367	CH3CH2-	$\bigcirc \dashv$	Н	н
YA1368	СНЗСН2-	$\bigcirc$ $\dashv$	н	н
YA1369	СНЗСН2-		н	н
YA1370	СНЗСН2-	<b>⊘</b> -₁	н	Н
YA1371	СНЗСН2-		н	н
YA1372	снзсн2-		Н	н
YA1373	СНЗСН2-		. н	н
YA1374	снзсн2-		н	н
YA1375	снзсн2-		н	Н
YA1376	снасн2-	<del></del>	н	Н
YA1377	снасн2-	- Cler4	н	Н
YA1378	СНЗСН2-	CI 	н	н
YA1379	СНЗСН2-		н	н
YA1380	снзсн2-	; <b>-{_}</b> -{	н	Н
YA1381	снзсн2-		н	н
YA1382	CH3CH2-C		н	Н
YA1383	снзсн2-		н	н
YA1384	СНЗСН2-		н	н
YA1385	снзсн2- В		н	н
YA1386	снзсн2- В	<b></b>	н	Н

No.	R1	R2	R3	R4
YA1387	снзсн2-	Br—(	н	н
YA1388	снзсн2-		н	Н
YA1389	СНЗСН2-		Н	н
YA1390	снзсн2-		Н	Н
YA1391	СНЗСН2-	\ <u>}</u>	н	н
YA1392	СНЗСН2-	H₃C ——	Н	Н
YA1393	СНЗСН2-	H₃C-⟨}{	Н	Н
YA1394	снзсн2-	1	н	Н
YA1395	снасн2-		Н	H
YA1396	СНЗСН2-	<u> </u>	н	Н
YA1397	СНЗСН2-	OH	н	Н.
YA1398	СНЗСН2-	HO	н	Н
YA1399	СНЗСН2-		н	Н
YA1400	снзсн2-	OCH₃ <—}	н	Н
YA1401	снзсн2-	H₃CO <u></u>	н	Н
YA1402	СНЗСН2-	H₃CO- <b>⟨_</b> }{	н	Н
YA1403	СНЗСН2-	H₃CO- <b>⟨_</b> }-{	. н	Н
YA1404	СНЗСН2-	٠ _	н	Н
YA1405	СНЗСН2-	OC <sub>2</sub> H <sub>5</sub>	н	Н
YA1406	СНЗСН2-	C₂H₅O △_>⊣	н	Н
YA1407	СНЗСН2-	C <sub>2</sub> H <sub>5</sub> O-{_}-	н	Н



No.	R1		<del></del> -	
NO.		R2	R3	R4
YA1408	СН3СН2-	'n∙C₃H <sub>7</sub> O-⟨ <u></u> _⟩−₁	н	н
YA1409	снзсн2-	љ€₄ℍ₀O-⟨⟨⟩ <mark>─</mark> ┤	н	Н
YA1410	СНЗСН2-	\/	н	н
YA1411	снзсн2-	O <sub>2</sub> N	н	Н
YA1412	СНЗСН2-		н	Н.
YA1413	СНЗСН2-	CN	н	н
YA1414	СНЗСН2-	NC ———	н	н
YA1415	СНЗСН2-		н	н
YA1416	СНЗСН2-	NH <sub>2</sub>	н	н .
YA1417	СНЗСН2-	H <sub>2</sub> N	н	Н
YA1418	СНЗСН2-		н	Н
YA1419	СНЗСН2-	NMe <sub>2</sub>	н	. н
YA1420	СНЗСН2-	Me <sub>2</sub> N	Н	н
YA1421	СНЗСН2-	Me <sub>2</sub> N-(){	н	н
YA1422	СНЗСН2-		н	н
YA1423	СНЗСН2-	(n-()	н	н
YA1424	СНЗСН2-	_N-{_}-	н	н
YA1425	CH3CH2-	O+\(\sigma\)	Н	Н
YA1426	СНЗСН2-		н	Н
YA1427	СНЗСН2-		Н	Н
YA1428	СНЗСН2-		н	Н



				·
No.	R1	R2	R3	R4
YA1429	СНЗСН2-		н	н
YA1430	СНЗСН2-	<b>€</b> i-€]-{	Н	Н
YA1431	СНЗСН2-	H3CN N-	Н	н
YA1432	СНЗСН2-	H <sub>2</sub> CN N-{}	Н	н
YA1433	СНЗСН2-	H3CN_N-{}-{	Н	н
YA1434	СНЗСН2-	[ <b>\</b> \ '	Н	н
YA1435	СНЗСН2-	OCH₃ F	н	Н
YA1436	СНЗСН2-	OCH3 F—Out	Н	Н
YA1437	СНЗСН2-		Н	Н
YA1438	СНЗСН2-		Н	Н
YA1439	СНЗСН2-	CH3-	н	СН3-
YA1440	СНЗСН2-	снзсн2-	н	снз-
YA1441	СНЗСН2-	<b>^</b> \	н	СН3-
YA1442	СНЗСН2-	Y	н	CH3-
YA1443	СНЗСН2-	<b>\\\</b>	Н	СН3-
YA1444	СНЗСН2-	人、	н	СН3-
YA1445	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	СН3-
YA1446	СН3СН2-	丫	н	CH3-
YA1447	снзсн2-	<b>~</b> ~√\	н	CH3-
YA1448	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	CH3-
YA1449	СНЗСН2-	×1	Н	СН3-

No.	. R1	R2	R3	D4
YA1450		入入	Н	R4 CH3-
YA1451	снзсн2-	~~~	Н .	CH3
YA1452	снзсн2-	L~~	н	CH3-
YA1453	снзсн2-	~~~\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н	снз-
YA1454	СНЗСН2-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	снз-
YA1455	СНЗСН2-	~~~```````````````````````````````````	Н	СН3-
YA1456	снзсн2-		Н	СН3-
YA1457	СН3СН2-		Н	СН3-
YA1458	СНЗСН2-		н	СН3-
YA1459	СНЗСН2-		н	CH3-
YA1460	СНЗСН2-	<b>&gt;</b> → .	Н	CH3-
YA1461	СНЗСН2-	$\Diamond$ -I	Н	CH3-
YA1462	СНЗСН2-	$\bigcirc \dashv$	Н	СН3-
YA1463	СНЗСН2-	$\bigcirc$ $\vdash$	Н	CH3-
YA1464	СНЗСН2-	$\bigcirc$ $\dashv$	н	СН3-
YA1465	СНЗСН2-		н	СН3
YA1466	СНЗСН2-	<b>△</b> -1	. н	СН3
YA1467	СНЗСН2-		н	снз-
YA1468	СНЗСН2-		н	СН3-
YA1469	снзсн2-		н	CH3-
YA1470	СНЗСН2-		Н	снз-

•	~ ~	

No.	Rt	R2	R3	R4
YA1471	СНЗСН2-	L (1)	н	CH3-
YA1472	СНЗСН2-	F-Ond	н	СН3-
YA1473	СНЗСН2-	CI	н	CH3-
YA1474	СНЗСН2-	CI	н	СН3-
YA1475	СНЗСН2-	c <del>{_</del> }	н	CH3-
YA1476	СНЗСН2-	c <del>(_&gt;-</del>	н	СН3-
YA1477	СНЗСН2-	<u> </u>	Н	CH3-
YA1478	СН3СН2-	(_)_{	н	снз-
YA1479	СНЗСН2-	Br.	Н	СН3-
YA1480	СНЗСН2-	Br—(	н	CH3-
YA1481	СНЗСН2-	Br- <b>⟨</b> }-{	н	CH3-
YA1482	СНЗСН2-	Br-{_}\m\{	н	.СН3-
YA1483	СНЗСН2-		н	CH3-
YA1484	СНЗСН2-	<u></u>	н	CH3-
YA1485	СНЗСН2-	<del></del>	н	СН3-
YA1486	СНЗСН2-	CH₃ ————————————————————————————————————	н	СН3-
YA1487	СНЗСН2-	H <sub>3</sub> C	. Н	СН3
YA1488	СНЗСН2-	H <sub>3</sub> C-{_}}-{	н	СН3-
YA1489	СНЗСН2-	C <sub>2</sub> H <sub>5</sub> -{_}-{	н .	СН3-
YA1490	СНЗСН2-		н	CH3-
YA1491	СНЗСН2-	n-C <sub>4</sub> H <sub>9</sub> -{}-{	н	СН3-

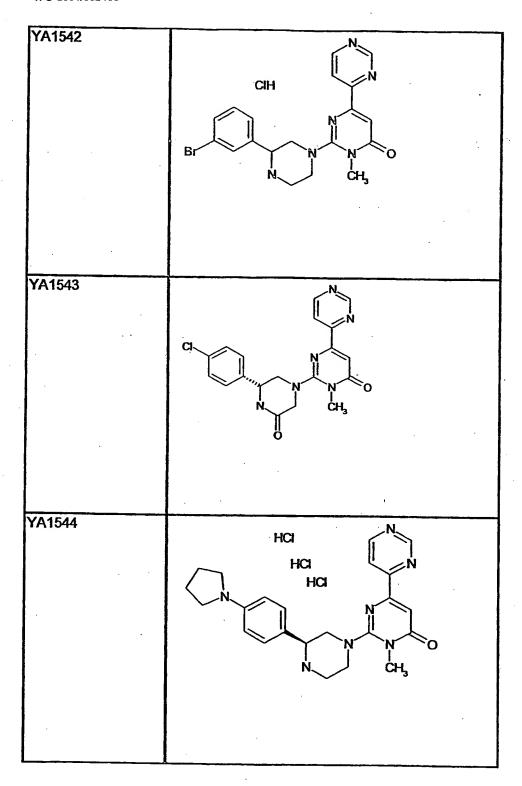
No.	D1	<del></del>		
140.	R1	OH R2	R3	R4
YA1492	снзсн2-		н	снз-
YA1493	СНЗСН2-	HO \	н	СН3-
YA1494	СНЗСН2-		н	СН3
YA1495		OCH₃  	Н	CH3-
YA1496	СНЗСН2-	H <sub>3</sub> CO	н	СН3-
YA1497	СНЗСН2-	H <sub>3</sub> CO-{_}_{	н	снэ-
YA1498	снзсн2-	H₃CO-{\_}{\}	н	СН3-
YA1499	СНЗСН2-		Н	СН3
YA1500	СНЗСН2-	OC <sub>2</sub> H <sub>5</sub>	Н	СН3-
YA1501	СНЗСН2-	<sup>2</sup> <sub>2</sub> H <sub>5</sub> O <del></del> <del></del> <del></del> <del></del> <del></del> <del></del> <del></del> <del></del> <del></del>	н	СН3-
YA1502	CH3CH2-	<sup>2</sup> 2H₅O-{_}_{}	н	СН3-
YA1503	снзсн2-	-C₃H <sub>7</sub> O-⟨_}{	н	СН3-
YA1504	CH3CH2-		н	СН3-
YA1505	СНЗСН2-	NO <sub>2</sub>	н	СН3-
YA1506	СНЗСН2-	Z <sub>2</sub> N	Н	CH3-
YA1507	снзсн2-	2N-()(	H	CH3-
YA1508	снзсн2-	CN	. н	CH3-
YA1509	СНЗСН2-		н	CH3-
YA1510	снзсн2-	<u> </u>	н	СН3-
YA1511	снзсн2-	NH <sub>2</sub>	н	CH3-
YA1512	CH3CH2-	,N	н	СН3-

			·	
No.	R1	R2	R3	R4
YA1513	снзсн2-		н	снз-
YA1514	снзсн2-		Н	СН3
YA1515	СНЗСН2-	Me <sub>2</sub> N	Н	CH3-
YA1516	снзсн2-	Me <sub>2</sub> N-{}-{	Н	CH3-
YA1517	СН3СН2-		н	CH3-
YA1518	СНЗСН2-		Н	CH3-
YA1519	СНЗСН2-		н	CH3-
YA1520	СНЗСН2-		н	СН3
YA1521	СНЗСН2-		Н	СН3-
YA1522	СНЗСН2-	()+( <u>)</u> -1	н	СН3
YA1523	СНЗСН2-		н	СН3-
YA1524	СНЗСН2-	○~	н	СН3-
YA1525	снзсн2-		Н	СН3-
YA1526	СНЗСН2-		Н	снз-
YA1527	СНЗСН2-	H3CN N-{_}	Н	СН3
YA1528	СНЗСН2-	H³CN_N-{}-{	н	снз-
YA1529	СНЗСН2-		Н	СН3-
YA1530	СНЗСН2-	<b>'</b> _∕ '	н	СН3-
YA1531	СНЗСН2-		Н	снз-
YA1532	СНЗСН2-		н	CH3-
YA1533	СНЗСН2-	M,	н	CH3-

No.	STRUCTURE	
YA1534		
	CH <sub>3</sub> N N N N N O CH <sub>3</sub> O CH <sub>3</sub>	
YA1535		
·	CIH CIH	
	CI N N CH <sub>3</sub>	
YA1536	N	
	CIH CIH N	
	CI N N CH <sub>3</sub>	
'A1537	N N	
	H <sub>3</sub> C CH <sub>3</sub>	

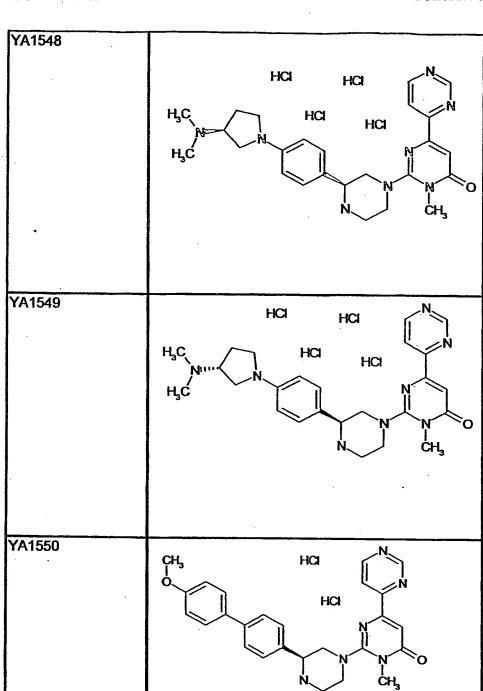
5/44500	
YA1538	OH CH <sub>3</sub>
YA1539	
	N N
·	H <sub>3</sub> C N CH <sub>3</sub>
YA1540	ĊН₃
	H <sub>3</sub> C O CH <sub>3</sub>
YA1541	CI N N CH <sub>3</sub>



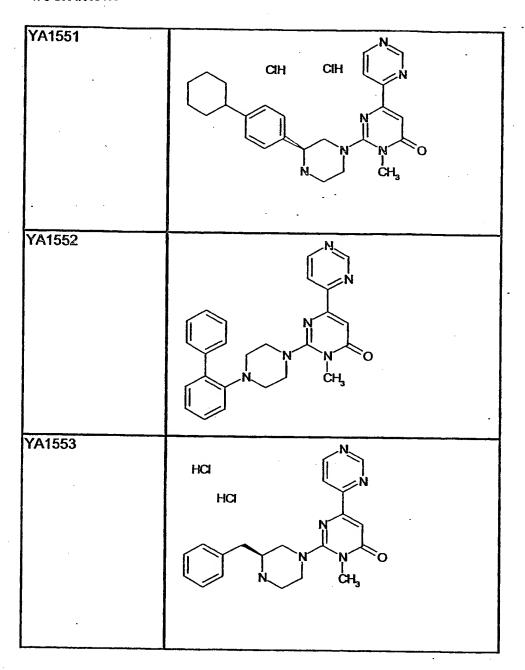




YA1545	Ha Ha Ha Ha A A A A A A A A A A A A A A
YA1546	HCI HCI HCI HCI CH <sub>3</sub>
YA1547	H <sub>3</sub> C

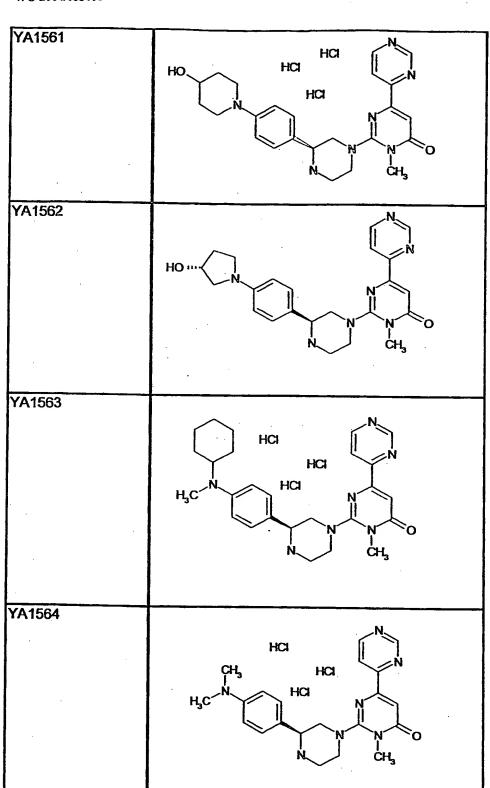




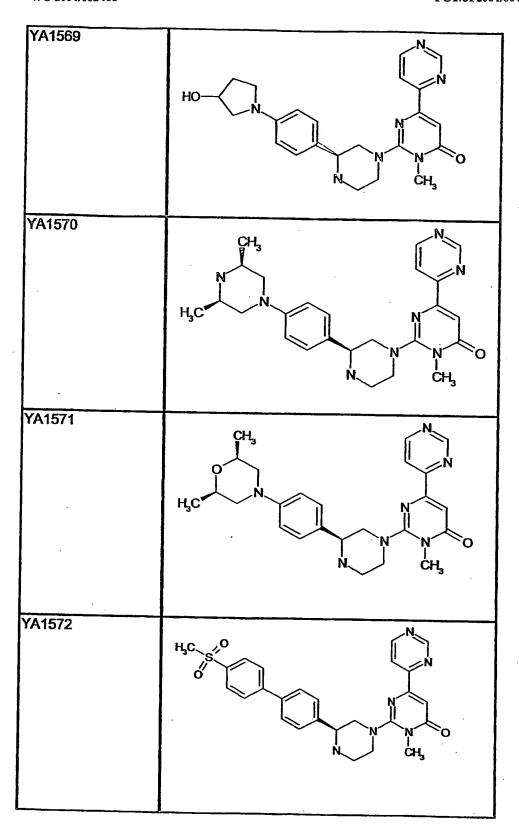


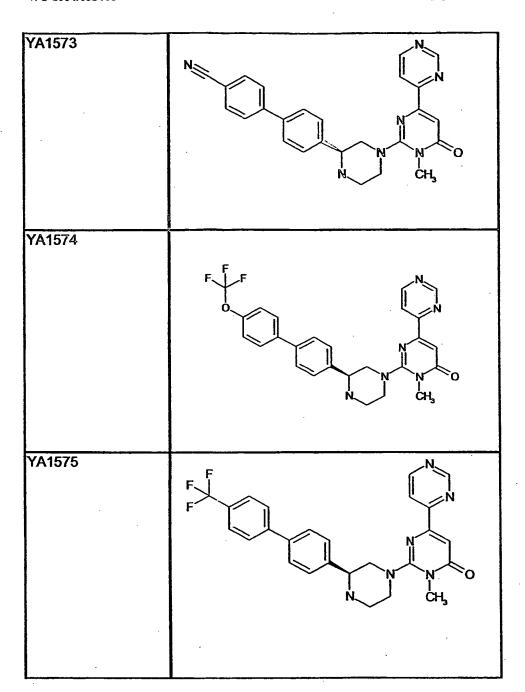
YA1554	
FA1354	Ha Ha N N N CH <sub>3</sub>
YA1555	HCI HCI N N N N CH <sub>3</sub>
YA1556	HCI HCI HCI NO CHIS
YA1557	HCI HCI HCI N N N N O CH <sub>3</sub>

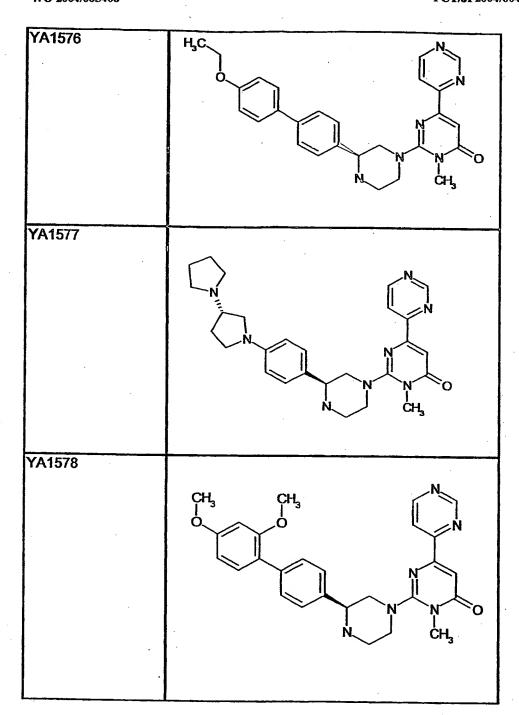
544.4550	
YA1558	H <sub>3</sub> C <sub>3</sub> O <sub>N</sub> O <sub>CH<sub>3</sub></sub> O <sub>CH<sub>3</sub>O<sub>CH<sub>3</sub></sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub></sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub></sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub>O<sub>CH<sub>3</sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub></sub>
YA1559	Ha Ha Ha Ha Ha N N N N N N N N N N N N N
YA1560	Ha H

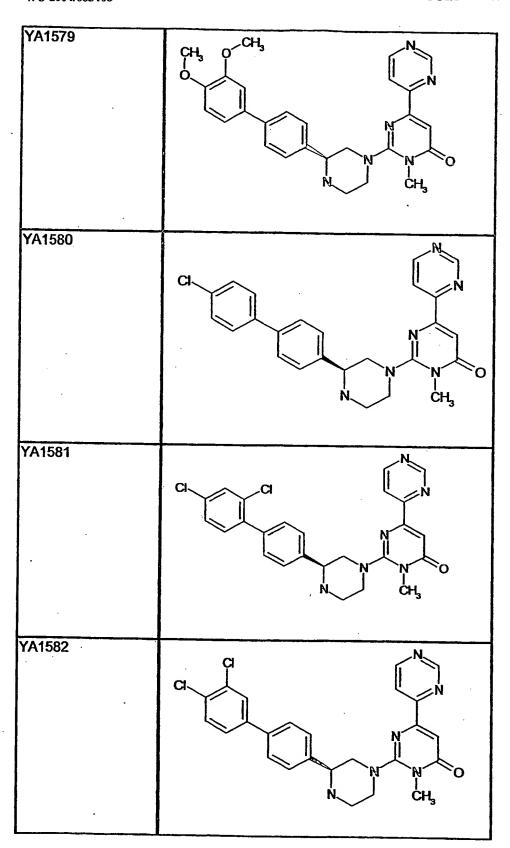


YA1565	
YA1566	CH <sub>3</sub> H <sub>3</sub> C N N N N N N N N N N N N N N N N N N N
YA1567	HO NO CH <sub>3</sub>
YA1568	N N CH <sub>3</sub>









224/1

D/A 4700	
YA1583	O N N N CH <sub>3</sub>
YA1584	H <sub>3</sub> C O N N N O CH <sub>3</sub>
YA1585	H <sub>3</sub> C O CH <sub>3</sub>
YA1586	CH <sub>3</sub> S N N N N CH <sub>3</sub> CH <sub>3</sub>

VA4507	T
YA1587	H <sub>3</sub> C N N N N N N N N N N N N N N N N N N N
YA1588	H <sub>3</sub> C H <sub>3</sub>
YA1589	H <sub>2</sub> N N CH <sub>3</sub>
YA1590	Br N O CH <sub>3</sub>

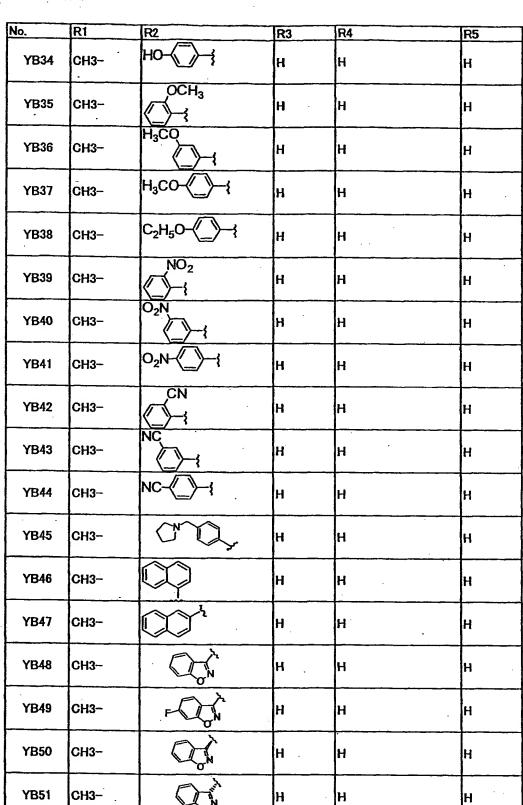
225/1





Table-4					
	•	Sh.			
		R <sub>3</sub> N N O R <sub>4</sub> R <sub>5</sub>	·		
No.	R1	R2	R3	R4	R5
YB1	снз-	СН3-	н	Н	н
YB2	СН3-	CH3CH2-	н	Н	н
YB3	снз-	<u> </u>	н	Н	Н
YB4	СН3-	Y	Н	Н	н
YB5	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	н	н .
YB6	СН3-	人、	Н	н	Н
YB7	СН3-	丫	Н	н	Н
YB8	снз-	^^\	Н	н .	н
YB9	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	н	Н
YB10	СН3	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Н	Н
YB11	СН3-	~~~\\	н	н	Н
YB12	СН3-	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н	Н	H
YB13	СН3-	Oor	н	н	Н
YB14	СН3-	Qui	Н	Н	Н
YB15	СН3-	Q	Н	Н	Н

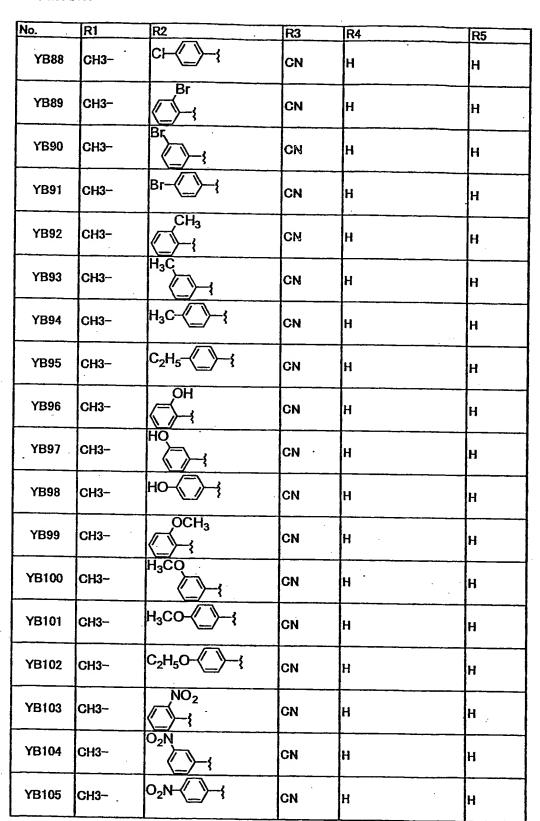
No.	R1	R2	R3	R4	R5
YB16	СН3-		Н	н	н
YB17	СН3-	Ď	н	н	н
YB18	СН3-	<b>◯</b> ~₁	Н	Н	н
YB19	снз-	F →	н	Н	н
YB20	СН3-		Н	Н	н
YB21	СН3-	F-{}-{	Н	Н	Н
YB22	СН3-	CI	н	Н	Н
YB23	СН3	CI	Н	н	н
YB24	СН3-	c <del></del>	н	Н	Н
YB25	СН3-	Br	н	Н	н
YB26	СН3-	Br.	н	Н	н
YB27	СН3-	Br—{	н	Н	н
YB28	СН3-	CH <sub>3</sub>	н	н .	н
YB29	СН3-	H <sub>3</sub> C	Н	Н	н
YB30	СН3-	H <sub>3</sub> C-{}-{	Н.	Н	Н
YB31	снз-	C <sub>2</sub> H <sub>5</sub> {	н	H	Н
YB32	СН3-	OH ⟨□⟩→	Н	н	н
YB33	СН3~ .	HO ————————————————————————————————————	Н	Н	Н



No.	R1	R2	102	D4	R5
140.	-   N		R3	R4	- Ra
YB52	снз-	<u></u>	ОН	н	н
YB53	СН3-	F	он	Н	н
YB54	CH3-	F	ОН	Н	н
YB55	СН3-		он	Н	н
YB56	СН3-	CI	он	н	н
YB57	СН3-	CI	он	н	н
YB58	снз-	C⊢{_}_{	ОН	н	н
YB59	СН3-	Br	ОН	Н	н
YB60	снз–	Br.	он	Н	н
YB61	снз-	Br—{{{}^{\!	он	Н	. Н
YB62	СН3-	CH₃	ОН	Н	Н
YB63	СН3-	H <sub>3</sub> C	он	Н	Н
YB64	СН3-	H₃C-⟨{	ОН	Н	Н
YB65	СН3-	C <sub>2</sub> H <sub>5</sub> -{	ОН	Н	н
YB66	снз-	ОН	ОН	Н	н
YB67	снз-	HO ————————————————————————————————————	ОН	н	Н
YB68	СН3-	HO-{}-{	он	Н	н
YB69	СН3-	OCH <sub>3</sub>	он	н	н



No.	R1	R2	R3	R4	R5
YB70	СН3-	H <sub>3</sub> CO	он	Н	Н
YB71	СН3-	H <sub>3</sub> CO-{\bigs\}-{	он	н	Н
YB72	СН3-	C <sub>2</sub> H <sub>5</sub> O-{	он	Н	Н
YB73	СН3-	NO <sub>2</sub>	он	н	н
YB74	СН3-	O <sub>2</sub> N	ОН	Н	н
YB75	СН3-	O <sub>2</sub> N-{	он	н	н
YB76	СН3-	CN	он	н	н
YB77	СН3-	NC	он	н	H
YB78	СН3-	NC-{	ОН	Н	н
YB79	СН3-	and,	ОН	Н	н
YB80	снз-		ОН	Н	Н
YB81	снз-	CC,	ОН	Н	Н
YB82	СН3-		CN	н	н
YB83	СН3-	F.	CN	н .	н
YB84	СН3-		CN	н	Н
YB85	СН3-	F-()	CN	Н	Н
YB86	снз-	CI	CN	н	Н
YB87	СН3-	CI	СИ	Н	н

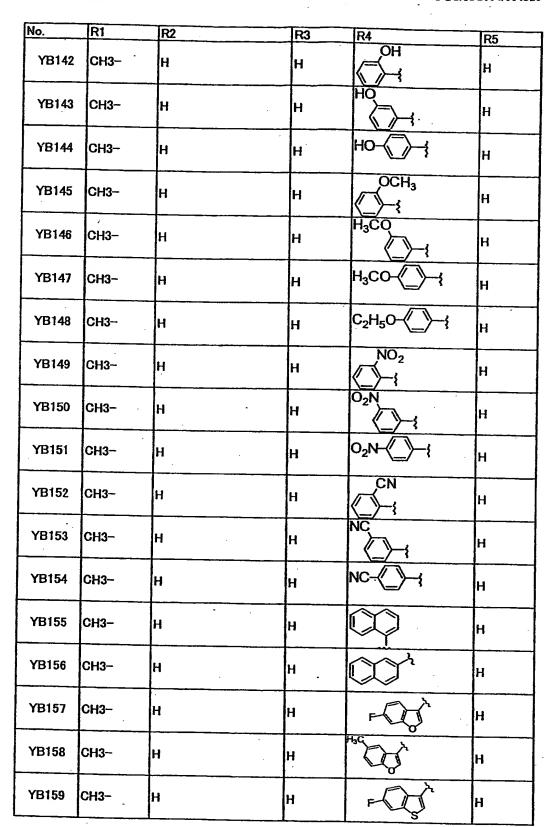


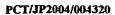


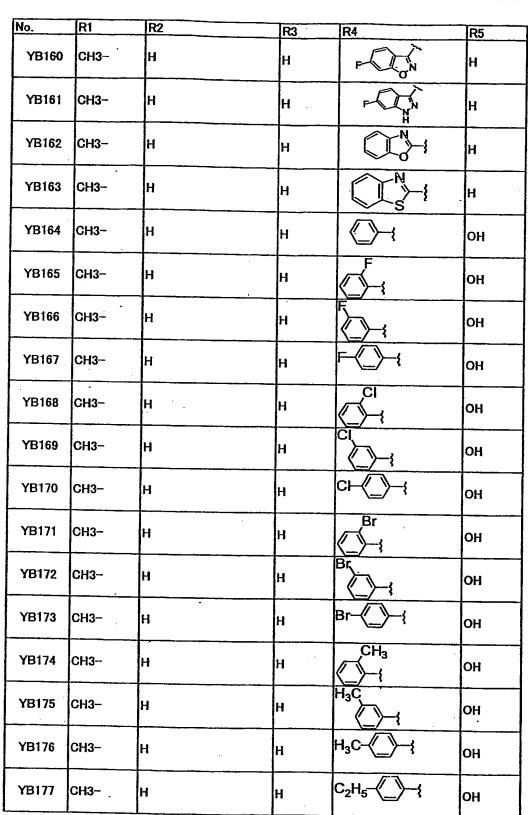
No.	R1	R2	R3	In4	1==
	T	CN	17.3	R4	R5
YB106	СН3-		CN	н	н
YB107	СН3-	NC	CN	Н	Н
YB108	СН3	NC-()-(	CN	н	н
YB109	СН3-	ChO,	CN	Н	н
YB110	снз-		СИ	Н	Н
YB111	снз-		CN	Н	н
YB112	снз–	н	Н	СН3-	Н
YB113	снз-	Н	н	СН3СН2-	Н
YB114	СН3-	Н	Н	<b>^</b> \	Н
YB115	СН3-	Н	Н	7	н
YB116	СН3-	Н	Н	<b>\\\\</b>	Н
YB117	СН3-	Н	н	人、	Н
YB118	СН3-	н	Н	大 <sup>'</sup> '	н
YB119	СН3-	Н	Н	<b>^</b> \\\\	н
YB120	СН3-	Н	н .	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	н
YB121	СН3-	н	Н	<b>\\\\</b>	Н
YB122	СН3-	Н	Н	<b>~</b> ~~	Н
YB123	СН3-	Н	Н	\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\\	Н



No.	R1	R2	R3	R4	R5
YB124	СН3-	Н	Н	Q	Н
YB125	СН3-	н	Н		н
YB126	СН3-	н -	н	Q	Н
YB127	СН3-	н	Н		Н
YB128	СН3-	Н	Н	F-4	н
YB129	СН3-	Н	Н	<b>-</b>	Н
YB130	СН3-	н	н	F-(){	Н
YB131	СН3-	н	н	CI	н
YB132	СН3-	Н	Н	CI	Н
YB133	СН3-	Н	н	C⊢(_){	Н
YB134	снз-	Н	Н	CI—	н
YB135	снз–	Н	H	Br	н
YB136	СН3-	н	Н	Br.	Н
YB137	СН3-	н	Н	Br—{_}	н
YB138	СН3-	Н	Н	CH₃	н
YB139	СН3-	н	н	H <sub>3</sub> C	Н
YB140	СН3-	н	Н	H <sub>3</sub> C-{{}{	Н
YB141	CH3-	Н	н	C <sub>2</sub> H <sub>5</sub> -{	Н

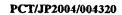






No.	R1	R2	Inc	104	
110.	1	1/2	R3	R4 OH	R5
YB178	СН3-	Н	Н		он
YB179	СН3-	Н	н	HO —	он
YB180	СН3-	н	Н	HO-{}-	он
YB181	СН3-	н	Н	OCH <sub>3</sub>	ОН
YB182	СН3-	н	Н	H₃CO —	ОН
YB183	СН3-	н	Н	H₃CO-⟨}	-{ ОН
YB184	СН3-	Н	н	C <sub>2</sub> H <sub>5</sub> O-	<sup>—{</sup> OH
YB185	СН3-	н	н	NO <sub>2</sub>	ОН
YB186	СН3-	н	Н	O <sub>2</sub> N	ОН
YB187	СН3-	н	н	O <sub>2</sub> N-(	ОН
YB188	СН3-	н	н	CN ←	ОН
YB189	СН3-	Н	Н	NC NC	ОН
YB190	СН3-	н	н	NC-{}-{	он
YB191	CH3-	Н	Н		он
YB192	СН3-	Н	Н	O)'	ОН
YB193	СН3-	н	Н		CN
YB194	CH3-	Н	Н	F	CN
YB195	СН3-	Н	н	F	CN

No.	R1	R2	R3	R4	R5
YB196	СН3-	Н	Н	F—()—{	CN
YB197	СН3-	Н	Н	CI	CN
YB198	СН3-	Н	Н	CI	CN
YB199	СН3-	Н	Н	CH	CN
YB200	СН3-	Н	Н	Br	CN
YB201	СН3-	Н	Н	Br.	CN
YB202	CH3-	н	Н	Br <b>-</b> ⟨{}-{	CN
YB203	СН3-	н	Н	CH₃	GN
YB204	СН3-	Н	Н	H <sub>3</sub> C —}	CN
YB205	снз-	Н	Н	H <sub>3</sub> C-	CN
YB206	СН3-	Н	н	C <sub>2</sub> H <sub>5</sub> -{}-{	CN
YB207	СН3-	н	Н	OH ⟨□}	CN
YB208	снз-	Н	Н	HO ————————————————————————————————————	CN
YB209	СН3-	н	Н	HO-{}-{	CN
YB210	СН3-	н	Н .	OCH <sub>3</sub>	CN
YB211	СН3-	н	Н	H₃CO —}	CN
YB212	СН3-	Н	н	H <sub>3</sub> CO-{	CN
YB213	СН3-	Н	Н	C <sub>2</sub> H <sub>5</sub> O-{}	CN



No.	R1	R2	R3	R4	Inc
YB214		Н	Н	NO <sub>2</sub>	R5 CN
YB215	CH3-	Н	Н	O <sub>2</sub> N	CN
YB216	СН3-	н	н	0214	CN
YB217	СН3-	Н	· H	CN CN	CN
YB218	снз-	н	н	NC NC	CN
YB219	СН3-	Н	н	NC-{}	CN
YB220	СН3-	H	Н		СИ
YB221	СН3-	Н	Н	CO'	CN
YB222	СН3-	Н	Н	<b>⟨</b> }-;	0
YB223	снз-	н	Н	F	0
YB224	СН3-	Н	Н	F;	٠ •
YB225	СН3-	Н	н	F-(){	0
YB226	СН3-	Н	н	CI →	<u></u>
YB227	СН3-	Н	н	CI	٠
YB228	СН3-	н	Н	CI—()—{	0
YB229	СН3-	Н	н	Br	<u></u>
YB230	СН3-	Н	Н	Br.	0
YB231	СН3-	Н	н	Br—	0

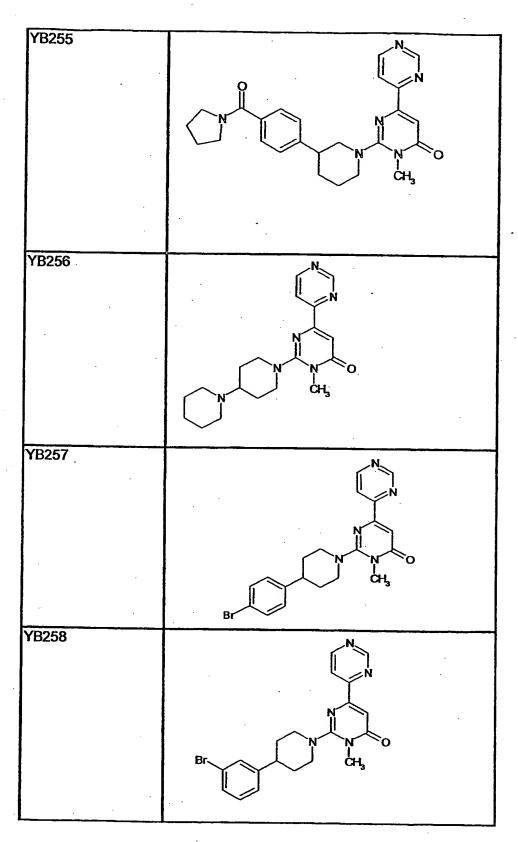
No.	R1	R2			
			R3	R4 CH <sub>3</sub>	R5
YB232	CH3-	Н	н		
YB233	СН3-	н	н	H <sub>3</sub> C	<u></u>
YB234	СН3-	н	н	H <sub>3</sub> C-{}-{	0
YB235	СН3-	н	н	C <sub>2</sub> H <sub>5</sub> —{	<u></u>
YB236	СН3-	Н	н	OH	0
YB237	СН3-	Н	Н	HO	0
YB238	СН3-	Н	Н	HO-{}-}	0
YB239	СН3-	Н	Н	OCH <sub>3</sub>	<u></u>
YB240	СН3-	Н	н	H <sub>3</sub> CO	0
YB241	СН3-	н	Н	H <sub>3</sub> CO-{}-{	0
YB242	СН3-	Н	н	C <sub>2</sub> H <sub>5</sub> O-{}-{	0
YB243	СН3-	Н	н	NO <sub>2</sub>	O.
YB244	СН3-	Н	н	O <sub>2</sub> N	0
YB245	СН3-	н	Н	O <sub>2</sub> N-{}-{	0
YB246	СН3-	н	Н	CN ←	<u></u>
YB247	CH3-	н	н	NC	Ů,
YB248	СН3-	Н	н	NC-	0
YB249	СН3-	Н	Н		<u></u>



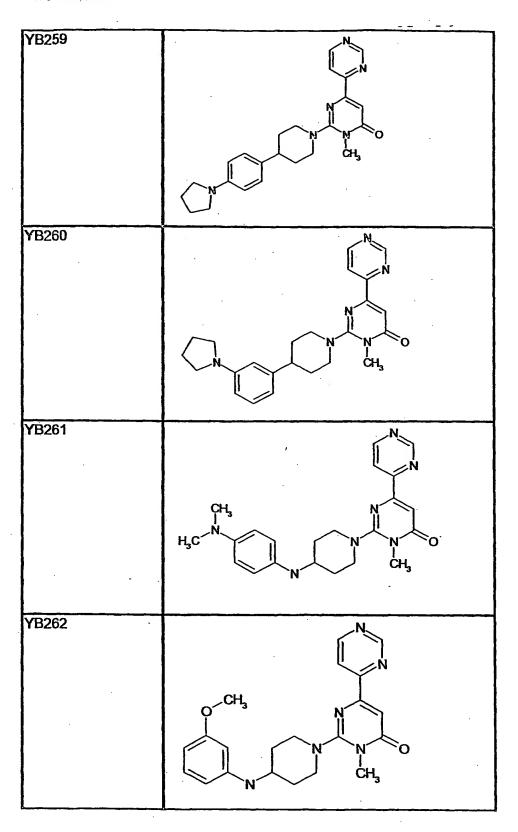
No.	R1	R2	R3	R4	R5
YB250	СН3-	н	Н		0



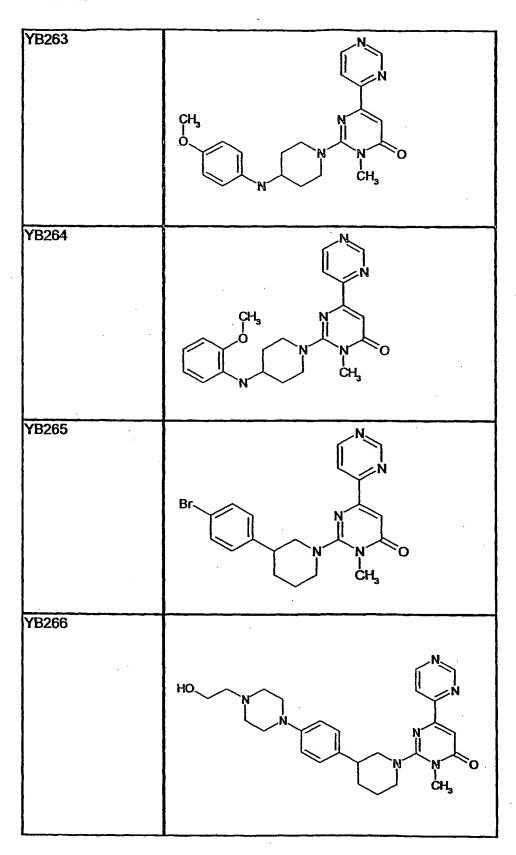
No.	STRUCTURE
YB251	o H o H,
YB252	CH <sub>3</sub>
YB253	S CH,
YB254	ON OH OH











242/1

YB272	O CH <sub>3</sub>
YB273	H <sub>3</sub> C N CH <sub>3</sub>
YB274	HO CH <sub>3</sub>
YB275	N CH <sub>3</sub>



# · ...

YB267	H <sub>3</sub> C N N O CH <sub>3</sub>
YB268	
YB269	N N CH <sub>3</sub>
YB270	H <sub>3</sub> C N N CH <sub>3</sub>
YB271	CH <sub>3</sub> H <sub>3</sub> C N N N N CH <sub>3</sub>





YB276	O N CH,
YB277	N N N CH <sub>3</sub>
YB278	CH <sub>3</sub>



Particularly preferred compounds of the present invention represented by formula (I) include:

2-(3-Phenylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
2-(3-(4-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
2-(3-(3-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
2-(3-(2-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
2-(3-(4-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
(S)-2-(3-(4-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;

(R)-2-(3-(4-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;

2-(3-(3-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(4-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(3-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(4-Methylphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(3-Methylphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Methylphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(4-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(3-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(4-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(3-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(2-Ethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(5-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3Hpyrimidin-4-one;





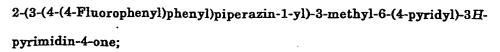
- $\hbox{2-(3-(4-Fluoro-3-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} H-10-(4-pyridyl)-3-(4-p$ pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3Hpyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3Hpyrimidin-4-one;
- pyrimidin-4-one;
- $\hbox{2-(3-(4-Chloro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} H$ pyrimidin-4-one;
- pyrimidin-4-one;
- pyrimidin-4-one;
- $\hbox{2-(3-(5-Bromo-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} H-pyrimid$ in-4-one;
- pyrimidin-4-one;
- $\hbox{2-(3-(2-Chloro-6-fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} H-10-(2-Chloro-6-fluorophenyl)$ pyrimidin-4-one;
- 2-(3-(2,4-Difluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4one;
- $\hbox{2-(3-(2,6-Difluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3$$H$-pyrimid in -4-pyrimid in$ one;
- 2-(3-(2,6-Dichlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4one;
- 2-(3-(2,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;





- 2-(3-(3,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,5-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,4-Difluoro-6-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(5-Cyano-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Cyano-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(1-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,3-Dihydrobenzofuran-7-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(Benzofuran-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; (S)-2-(3-(Benzofuran-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-4-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-5-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Phenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;





- 2-(3-(4-(4-Methoxyphenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(4\hbox{-}(2\hbox{-Methoxyphenyl}) phenyl) piperazin-1-yl)-3\hbox{-methyl-}6\hbox{-}(4\hbox{-pyridyl})-3H-pyrimidin-4-one;$
- 2-(3-(4-(Morpholin-4-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(4-Methylpiperazin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Phenylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzoylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-(1,2-Benzisothiazol-3-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Methyl-3-phenylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Acetyl-3-(4-fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzyl-3-(4-fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzyl-3-(ethoxycarbonyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;



- $\hbox{2-(4-methyl-3-(1-naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3$$H$-pyrimidin-4-methyl-6$ one;
- $2\hbox{-}(5,5\hbox{-}Dimethyl-3\hbox{-}(2\hbox{-}methoxyphenyl)piperazin-1-yl)-3\hbox{-}methyl-6\hbox{-}(4\hbox{-}pyridyl)-3H-10-yl)-3-piperazin-1$ pyrimidin-4-one;
- 2-(3-Phenylpiperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- $\hbox{2-(3-(2-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3} \textit{H-pyrimidin-4-one};$
- 2-(3-(4-Chlorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Bromophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Methoxyphenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(3\hbox{-}Methoxyphenyl) piperidin-1-yl)-3\hbox{-}methyl-6\hbox{-}(4\hbox{-}pyridyl)-3\textit{H-}pyrimidin-4\hbox{-}one;$
- 2-(3-(2-Methoxyphenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- pyrimidin-4-one;
- (S)-2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-Hydroxy-3-phenylpiperidin-1-yl)-3-methyl-6-(4-pyridyl)-3} \textit{H-pyrimidin-4-one};$
- $\hbox{2-(3-Phenylpiperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3$$H$-pyrimidin-4-one;}$
- 2-(3-(4-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(3\hbox{-}Fluorophenyl) piperazin-1-yl)-3-methyl-6\hbox{-}(4\hbox{-}pyrimidyl)-3$H-pyrimidin-4-one;$
- 2-(3-(2-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(4\hbox{-}\mathrm{Chlorophenyl}) \texttt{piperazin-1-yl})\hbox{-}3\hbox{-}\mathrm{methyl-6-}(4\hbox{-}\mathrm{pyrimidyl})\hbox{-}3\textit{H-pyrimidin-4-}$
- one;
- 2-(3-(3-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;

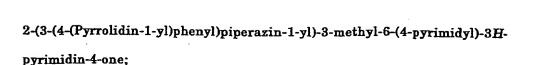




- 2-(3-(2-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(4-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(3-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(2-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(4-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(3-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(2-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(4-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(3-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-
- 2-(3-(2-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-
- 2-(3-(2-Ethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4one;
- 2-(3-(6-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3Hpyrimidin-4-one;
- 2-(3-(5-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3Hpyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3Hpyrimidin-4-one;



- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Chloro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(5-Bromo-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dichlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3*H*-pyrimidin-4-one;
- 2-(3-(2,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,5-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,4-Difluoro-6-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(1-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(2-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,3-Dihydrobenzofuran-7-yl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(Benzofuran-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;



- 2-(3-(2-methoxy-4-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one;
- 2-(3-(2-methoxy-5-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Phenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(4-Fluorophenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(4-Methoxyphenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(2-Methoxyphenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Morpholin-4-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(4-Methylpiperazin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(4-Acetyl-3-(4-fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(4-Benzyl-3-(4-fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;





2-(4-(4-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one; 2-(4-Cyano-4-phenylpiperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one; 2-(4-(6-Fluorobenofuran-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one;

- 2-(3-(Benzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(Benzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(Benzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(6-Fluorobenzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(4-(6-Fluorobenzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(4-(5-Methylbenzofuran-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; and
- 2-(4-(6-Fluorobenzothiophene-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one.

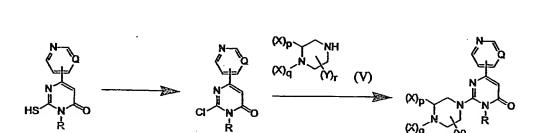
Salts of the aforementioned preferred compound, and solvates or hydrates of the aforementioned compounds and salts thereof are also preferred.

The 3-substituted-4-pyrimidone compounds represented by the aforementioned formula (I) can be prepared, for example, according to the method explained below.

(II)



(III)



(IV)

(In the above scheme, definitions of Q, R, X and Y are the same as those already described.)

The 2-thiopyrimidone represented by the above formula (III) is prepared easily by a modification of the method described in EP 354,179. The reaction may be carried out in the presence of a base such as sodium hydroxide, potassium hydroxide, sodium methoxide, sodium ethoxide, potassium tert-butoxide, sodium carbonate, sodium hydrogencarbonate, potassium carbonate, triethylamine, diisopropylethylamine, and 1,8-diazabicyclo[5,4,0]undec-7-en for 1 to 100 hours at a suitable temperature ranging from 0  $^{\circ}$  to 200  $^{\circ}$  under nitrogen or argon atmosphere or under ordinary air to afford the desired compound (III). Examples of a solvent for the reactions include, for example, alcoholic solvent such as methanol, ethanol, 1-propanol, isopropanol, tert-butanol, ethylene glycol, propylene glycol; etheric solvents such as diethyl ether, tert-butyl methyl ether, tetrahydrofuran, isopropyl ether; hydrocarbonic solvents such as benzene, toluene, xylene; halogenated hydrocarbonic solvents such as dichloromethane, chloroform, dichloroethane; aprotic polar solvents such as formamide, N,N-dimethylformamide, N,N-dimethylacetamide, N-methylpyrrolidone, dimethyl sulfoxide, sulfolane, hexamethylphosphoric triamide, water and the like. Generally, a single solvent or a mixture of two or more solvents may be used so as to be suitable to a base used.

Then the 2-thiopyrimidone derivative (III) is transformed into the 2-chloropyrimidone (IV) by a chlorinating agent. The reaction time and temperature depend on the chlorinating agent used. Examples of a chlorinating agent for the reactions include, for example, thionyl chloride, thionyl chloride and





dimethylformamide, phosphorus oxychloride, phosphorus oxychloride and dimethylformamide, oxalyl chloride, phosphorous oxychloride and dimethylformamide, and phosphorus pentachloride.

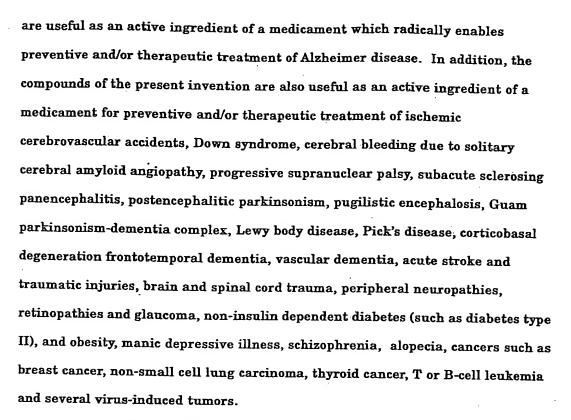
The amine represented by the above formula (V) may be prepared by a modification of the method described in Japanese Patent Unexamined Publication [Kokai] No. 52-139085/1977 or according to well-known methods of one skilled in the art.

Then the chloride derivative (IV) is allowed to react with the amine (V) or salts thereof in the presence of a base such as sodium hydroxide, potassium hydroxide, sodium methoxide, sodium ethoxide, sodium carbonate, sodium hydrogencarbonate, potassium carbonate, triethylamine, diisopropylethylamine, and 1,8-diazabicyclo[5,4,0]undec-7-en for 0.1 to 100 hours at a suitable temperature ranging from 0 °C to 200 °C under nitrogen or argon atmosphere or under ordinary air to afford the desired compound (II).

Examples of a solvent for the reactions include, for example, alcoholic solvent such as methanol, ethanol, 1-propanol, isopropanol, tert-butanol, ethylene glycol, propylene glycol; etheric solvents such as diethyl ether, tert-butyl methyl ether, tetrahydrofuran, isopropyl ether; hydrocarbonic solvents such as benzene, toluene, xylene; halogenated hydrocarbonic solvents such as dichloromethane, chloroform, dichloroethane; aprotic polar solvents such as formamide, N,N-dimethylformamide, N,N-dimethylacetamide, N-methylpyrrolidone, dimethyl sulfoxide, sulfolane, hexamethylphosphoric triamide, water and the like. Generally, a single solvent or a mixture of two or more solvents may be used so as to be suitable to a base used.

The compounds of the present invention have inhibitory activity against TPK1, and they inhibit TPK1 activity in neurodegenerative diseases like Alzheimer disease, thereby suppress the neurotoxicity of A $\beta$  and the formation of PHF and inhibit the nerve cell death. Accordingly, the compounds of the present invention





As the active ingredient of the medicament of the present invention, a substance may be used which is selected from the group consisting of the compound represented by the aforementioned formula (I) and pharmacologically acceptable salts thereof, and solvates thereof and hydrates thereof. The substance, per se, may be administered as the medicament of the present invention, however, it is desirable to administer the medicament in a form of a pharmaceutical composition which comprises the aforementioned substance as an active ingredient and one or more of pharmaceutical additives. As the active ingredient of the medicament of the present invention, two or more of the aforementioned substance may be used in combination. The above pharmaceutical composition may be supplemented with an active ingredient of other medicament for the treatment of, for example, Alzheimer disease, vascular dementia, acute stroke and traumatic injuries, brain and spinal cord trauma, peripheral neuropathies, retinopathies and glaucoma, non-insulin dependent diabetes (such as diabetes type II), and obesity, manic depressive illness,





schizophrenia, alopecia, cancers such as breast cancer, non-small cell lung carcinoma, thyroid cancer, T or B-cell leukemia and several virus-induced tumors.

A type of the pharmaceutical composition is not particularly limited, and the composition may be provided as any formulation for oral or parenteral administration. For example, the pharmaceutical composition may be formulated, for example, in the form of pharmaceutical compositions for oral administration such as granules, fine granules, powders, hard capsules, soft capsules, syrups, emulsions, suspensions, solutions and the like, or in the form of pharmaceutical compositions for parenteral administrations such as injections for intravenous, intramuscular, or subcutaneous administration, drip infusions, transdermal preparations, transmucosal preparations, nasal drops, inhalants, suppositories and the like. Injections or drip infusions may be prepared as powdery preparations such as in the form of lyophilized preparations, and may be used by dissolving just before use in an appropriate aqueous medium such as physiological saline.

Sustained-release preparations such as those coated with a polymer may be directly administered intracerebrally.

Types of pharmaceutical additives used for the manufacture of the pharmaceutical composition, content rations of the pharmaceutical additives relative to the active ingredient, and methods for preparing the pharmaceutical composition may be appropriately chosen by those skilled in the art. Inorganic or organic substances, or solid or liquid substances may be used as pharmaceutical additives. Generally, the pharmaceutical additives may be incorporated in a ratio ranging from 1% by weight to 90% by weight based on the weight of an active ingredient.

Examples of excipients used for the preparation of solid pharmaceutical compositions include, for example, lactose, sucrose, starch, talc, cellulose, dextrin, kaolin, calcium carbonate and the like. For the preparation of liquid compositions for oral administration, a conventional inert diluent such as water or a vegetable oil





may be used. The liquid composition may contain, in addition to the inert diluent, auxiliaries such as moistening agents, suspension aids, sweeteners, aromatics, colorants, and preservatives. The liquid composition may be filled in capsules made of an absorbable material such as gelatin. Examples of solvents or suspension mediums used for the preparation of compositions for parenteral administration, e.g. injections, suppositories, include water, propylene glycol, polyethylene glycol, benzyl alcohol, ethyl oleate, lecithin and the like. Examples of base materials used for suppositories include, for example, cacao butter, emulsified cacao butter, lauric lipid, witepsol.

Dose and frequency of administration of the medicament of the present invention are not particularly limited, and they may be appropriately chosen depending on conditions such as a purpose of preventive and/or therapeutic treatment, a type of a disease, the body weight or age of a patient, severity of a disease and the like. Generally, a daily dose for oral administration to an adult may be 0.01 to 1,000 mg (the weight of an active ingredient), and the dose may be administered once a day or several times a day as divided portions, or once in several days. When the medicament is used as an injection, administrations may preferably be performed continuously or intermittently in a daily dose of 0.001 to 100 mg (the weight of an active ingredient) to an adult.

## Examples

The present invention will be explained more specifically with reference to examples. However, the scope of the present invention is not limited to the following examples. The compound numbers in the examples correspond to those in the table above.

Reference Example 1: Synthesis of 2-mercapto-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one





A solution of ethyl 3-oxo-3-(4-pyridyl)propionate (29.0 g, 150 mmol), N-methyl thiourea (40.6 g, 450 mmol) and 1,8-diazabicyclo[5,4,0]-7-undecene (22.4 ml, 150 mmol) was refluxed for 4 hours and the solution of methanesulfonic acid (14.4 g, 150 mmol) in water (50 ml) was added after cooling by ice-water. The precipitate was washed with water, filtered and dried to give the title compound (23.7 g, 72%).

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>)  $\delta$ : 3.58(s, 3H), 6.40(s, 1H), 7.72(dd, J=1.8, 4.5Hz, 2H), 8.73(dd, J=1.5, 4.8Hz, 2H), 12.92(brd, 1H).

Reference Example 2: Synthesis of 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one

Phosphorous oxychloride (26.11g, 170 mmol) was added to dimethylformamide(180 ml) and stirred 20 min. 2-Mercapto-3-methyl-6-(4-pyridyl)-pyrimidine-4-one (24.15 g, 110 mmol) was added to the solution and stirred 5 min and then stirred at 70°C for 2 hours. Ethyl acetate (630 ml) was added to the ice-cooled solution and precipitate was collected by filtration after stirring for 20 minutes. After drying, the precipitate was dissolved in water (400 ml) and pH was adjusted to 10 by using aqueous sodium hydroxide. The precipitate was washed with water, filtered and dried to give the title compound (18.82 g, 77%).

1H-NMR (CDCl<sub>8</sub>)  $\delta$ : 3.72(s, 3H), 6.90(s, 1H), 7.78(dd, J=1.7, 4.5Hz, 2H), 8.75(dd, J=1.6, 4.5Hz, 2H).

Reference Example 3: Synthesis of 2-mercapto-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one

A solution of ethyl 3-oxo-3-(4-pyrimidyl)propionate (34.1 g, 176 mmol), N-methyl thiourea (47.5 g, 527 mmol) and 1,8-diazabicyclo[5,4,0]-7-undecene (26.3 ml, 176 mmol) in ethanol (340 ml) was refluxed for 2 hours and the solution of methanesulfonic acid (16.9 g, 176 mmol) in water (70 ml) was added after cooling by

ice-water. The precipitate was washed with water, filtered and dried to give the title compound (30.2 g, 78%).

<sup>1</sup>H-NMR (DMSO-d<sub>6</sub>)  $\delta$ : 3.56(s, 3H), 6.88(s, 1H), 8.24(dd, J=1.2, 5.4 Hz, 2H), 9.05 (dd, J=5.4 Hz, 1H), 11.94(s, 1H).

Reference Example 4: Synthesis of 2-chloro-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one

Phosphorous oxychloride (4.60 g, 30 mmol) was added to dimethylformamide(32 ml) and stirred for 20 min at 0°C. 2-Mercapto-3-methyl-6-(4-pyrimidyl)-3H-pyrimidine-4-one(4.40 g, 20 mmol) was added to the solution and stirred 5 min and then stirred at 70°C for 2 hours. The reaction mixture was poured into ice water, neutralized by solid potassium carbonate, and extracted with ethyl acetate. The organic layer was washed with brine, dried over sodium sulfate, and evaporated under reduced pressure. Purification of the residue by silica gel chromatography (ethyl acetate) gave the title compound (1.20 g, 27%).

1H-NMR (CDCls)  $\delta$ : 3.74(s, 3H), 7.56(s, 1H), 8.18(d, J=5.1 Hz, 1H), 8.92(d, J=5.1 Hz, 1H), 9.30(s, 1H).

MS[M+H]+: 223.

Example 1: Synthesis of 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-3*H*-pyrimidin-4-one dihydrochloride (No. XA468)

A solution of 2-bromo-5-fluoroanisole (11.8 g, 57.6 mmol) in tetrahydrofuran (60 ml) was dropped into the magnesium (1.40 g, 57.6 mmol) in refluxed tetrahydrofuran (32 ml) containing small amount of 1,2-dibromoethane and refluxed for 15 min. After addition of tetrehydrofuran (50 ml), the solution was cooled to -78 °C and diethyl oxalate (7.41 g, 50.7 mmol) in diethyl ether (50 ml) was dropped into the solution. After stirring at same temperature for 30 min, the solution was warmed to -10°C and 1N aqueous hydrogen chloride (50 ml) and water



(6.80g, 59%)



were added. Organic layer was extracted with diethyl ether, washed with brine and dried over magnesium sulfate. After removal of the solvent under reduced pressure, purification of the residue by silica gel column chromatography (eluent: hexane/ethyl acetate = 5/2) gave ethyl 2-(4-fluoro-2-methoxyphenyl)-2-oxoacetate

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 1.40(3H, t, J=7.1 Hz),3.87(3H, s), 4.89(2H, q, J=7.1Hz), 6.68(1H, d, J=10.5 Hz), 6.77-6.81(1H, m), 7.91-7.95(1H, m).

Ethylenediamine (0.60 g, 10.0 mmol) was added to a solution of ethyl 2-(4-fluoro-2-methoxyphenyl)-2-oxoacetate (2.26 g, 10.0 mmol) in ethanol(30 ml) and refluxed 4 hr. After removal of the solvent under reduced pressure, residue was washed with ethanol-diethyl ether to give 5,6-dihydro-3-(4-fluoro-2-methoxyphenyl)pyrazinone (1.76 g, 79%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 3.50-3.56 (2H, m), 3.81 (3H, s), 3.88-3.92 (2H, m), 6.65(1H, d, J=11.0 Hz), 6.70-6.76 (1H, m), 6.89(1H, bs), 7.36-7.40(1H, m).

5,6-Dihydro-3-(4-fluoro-2-methoxyphenyl)pyrazinone was added to the solution of lithium aluminium hydride (0.46 g, 12 mmol) in diethyl ether (25 ml) and refluxed for 6 hr. Water (0.48 ml), 15% sodium hydroxide solution (0.48 ml) and again water (1.21 ml) were added to the ice-cooled solution and the precipitate was filtered and washed with dichloromethane. Combined organic layer was evaporated to give 2-(4-fluoro-2-methoxyphenyl)piperazine (0.83 g, 99%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>) δ: 2.02(2H, s), 2.57-2.63 (1H, m), 2.80-2.89 (1H, m), 2.92-2.99 (2H, m), 3.06-3.12 (2H, m), 3.80(3H, s), 4.06 (1H, d, J=10.0 Hz), 6.56-6.65 (2H, m), 7.40 (1H, t, J=7.8 Hz).

2-Chloro-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (222 mg, 1.0 mmol) was added to an ice-cooled solution of 2-(4-fluoro-2-methoxyphenyl)piperazine (210 mg, 1.0 mmol), triethylamine (0.15 ml, 1.1 mmol) in N,N-dimethylformamide (10 ml) and stirred at that temperature for 1 hr and then at room temperature for 2 hr.

Next day, reaction was quenched by ice-water and the filtrate was washed with



water and dried to give 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (246 mg, 62%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 2.89-2.96 (1H, m), 3.19-3.31 (3H, m), 3.59 (3H, s), 3.62-3.74 (2H, m), 3.85 (3H, s), 4.39-4.44 (1H, m), 6.63-6.71 (2H, m), 6.67 (1H, s), 7.51-7.55 (1H, m), 7.81 (2H, dd, J=1.7, 4.6 Hz), 8.71 (2H, dd, J=1.7, 4.6 Hz).

4N Hydrogen chloride in 1,4-dioxane (0.4 ml) was added to the solution of 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (217 mg, 0.6 mmol) in dichloromethane (5 ml) and stirred for 15 min. After addition of diethyl ether, filtration and wash with diethyl ether and dryness gave the title compound (260 mg, quant.).

Example 2: Synthesis of 2-(2-(4-methoxyphenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one dihydrochloride (No. XA393)

Dimethylslufoxide (50 ml) solution of 4-methyoxyphenacylbromide (9.94 g, 43.4 mmol) and water (1.6 ml, 88.8 mmol) were stirred at 50°C for 2.5 hr. Water was added and the solution was extracted with ethyl acetate 3 times and washed with brine and then dried over sodium sulfate. Removal of the solvent gave 4-methoxyphenylglyoxal (8.30 g, quant.).

<sup>1</sup>H-NMR (DMSO)  $\delta$ : 3.84 (3H, s), 6.60-6.69 (1H, m), 7.04 (2H, d, J=8.8 Hz), 8.05 (2H, d, J=9.1 Hz).

Methanol (5 ml) solution of ethylenediamine (3.74 g, 62.29 mmol) was added to the ice-cooled solution of 4-methoxyphenylglyoxal (8.30 g, 45.5 mmol) in methanol (100 ml) and tetrahydrofuran (50 ml) and stirred for 10 min. After cooling to 0°C, sodium tetrahydroborate (6.14 g, 162.2 mmol) and additional methanol (50 ml) was added and stirred overnight. After removal of the solvent, aqueous sodium hydroxide was added and was extracted with dichloromethane three times and washed with brine and dried over sodium sulfate. After removal of the solvent, purification of the residue by silica gel column chromatography (eluent;

dichloromethane/ethanol/diethylamine = 20/2/1) gave 2-(4-methoxypheny)-piperazine (3.96 g, 45%).

<sup>1</sup>H-NMR (CDCl<sub>s</sub>) δ: 2.69(1H, dd, J=10.3, 11.9 Hz), 2.80-3.01(4H, m), 3.07-3.11 (1H, m), 3.68-3.73(1H, m), 3.79(3H, s), 6.84-6.88 (2H, m), 7.27-7.32 (2H, m).

A solution of triethylamine (697 mg, 6.9 mmol), 2-(4-methoxyphenyl)piperazine (430 mg, tetrahydrofuran (10 ml) was stirred at room temperature for 30
min and at 50°C for 3 hr. Solvent was removed under reduced pressure, and 1N
aqueous sodium hydroxide solution was added to the residue and extracted by
dichloromethane three times and washed with brine and dried over sodium sulfate.
After removal of the solvent under reduced pressure, the residue was purified by
silica gel column chromatography (eluent; dichloromethane/ethanol = 10/1) to give
2-(2-(4-methoxyphenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one
(594 mg, 76%)

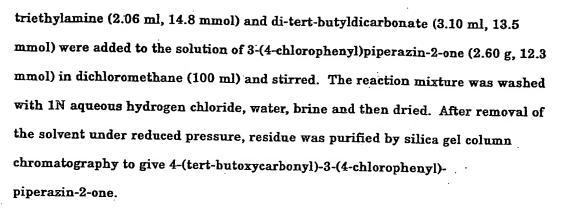
<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 3.02 (1H, dd, J=10.8, 12.7 Hz), 3.18-3.25 (3H, m), 3.55 (3H, s), 3.57-3.67 (2H, m), 3..82 (3H, s), 3.98(1H, dd, J=2.7, 10.8 Hz), 6.67 (1H, s), 6.92 (2H, d, J=8.7 Hz), 7.37 (2H, d, J=8.7 Hz), 7.80 (2H, d, J=6.0 Hz), 8.71 (2H, d, J=6.0 Hz).

4N Hydrogen chloride in ethyl acetate (5 ml) was added to the solution of 2-(2-(4-methoxyphenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (594 mg, 1.6 mmol) in dichloromethane (5 ml) and stirred for 1 hr. Wash with ethyl acetate after removal of the solvent and dryness gave the title compound (683 mg, 96%).

Example 3: Synthesis of 2-(2-(4-chlorophenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one hydrochloride (No. XA371)

Mixture of methyl (4-chlorophenyl)acetate (5.10 g, 27.6 mmol) and N-bromosuccinimide (5.16 g, 29 mmol) in carbon tetrachloride was treated by Hg lamp. After filtration, solvent was removed under reduced pressure and the residue was dissolved in methanol. Ethylenediamine (2.03 ml, 30.4 mmol) and





<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 1.44 (9H, s), 3.21-3.32 (2H, m), 3.48 (1H, m), 4.04 (1H, brs), 5.66 (1H, brs), 7.10 (1H, brs), 7.30-7.38 (4H, m).

Solution of 4-(tert-butoxycarbonyl)-3-(4-chlorophenyl)-piperazin-2-one (500 mg, 1.6 mmol) and acetic acid (929  $\mu$ l, 16 mmol) were added to a refluxed solution of sodium borohydride (608 mg, 16 mmol) in 1,4-dioxane (5 ml) and reflux was continued. The reaction was quenched by water and extracted with dichloromethane and washed with brine and dried. After removal of the solvent, residue was purified by silica gel column chromatography to give 4-(tert-butoxycarbonyl)-3-(4-chlorophenyl)piperazine (330 mg, 69%). <sup>1</sup>H-NMR (CDCl<sub>8</sub>)  $\delta$ : 1.46(9H, s), 2.76-2.99(3H, m), 3.13(1H, dd, J=13.0 Hz, 4.3 Hz), 3.45-3.49(2H, m), 3.92(1H, m), 5.15(1H, s), 7.27-7.33(4H, m).

A solution of 4-(tert-butoxycarbonyl)-3-(4-chlorophenyl)piperazine (330 mg, 1.1 mmol), 2-chloro-3-methyl-6-(4-pyridyl)pyrimidin-4-one (246 mg, 1.1 mmol) and triethylamine (170  $\mu$ l, 1.22 mmol) in tetrahydrofuran were refluxed. Usual workup and purification by silica gel column chromatography gave 2-(1-(tert-butoxy-carbonyl)-2-(4-chlorophenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (500 mg, 93%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 1.45(9H, s), 3.09(1H, m), 3,35(3H, s), 3.40-3.63(4H, m), 3.96-4.19(2H, m), 5.43(1H, s), 6.68(1H, s), 7.23(2H, d, J=8.3 Hz), 7.32(2H, d, J=8.3 Hz), 7.78(2H, d, J=5.9 Hz), 8.72(2H, d, J=5.9 Hz).

4N Hydrogen chloride in ethyl acetate was added to the solution of





2-(1-(tert-butoxycarbonyl)-2-(4-chlorophenyl)-piperazine-4-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (500 mg, 1.0 mmol) in ethyl acetate and stirred. Filtration and successive dryness gave the title compound (373mg, 79%).

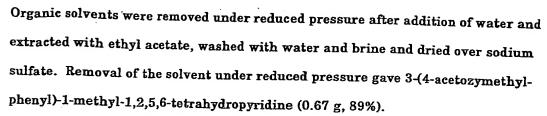
Example 4: Synthesis of 3-methyl-2-(3-(4-((1-pyrrolidinyl)methyl)phenyl)piperidine
-1-yl)-6-(4-pyridyl)pyrimidin-4-one fumarate (No. XB43)

Tetrakis(triphenylphosphine)palladium (0.65 g, 0.56 mmol),
4-formylphenylboric acid (2.81 g, 18.7 mmol), 2M aqueous sodium carbonate (18.7 ml, 37.4 mmol) and ethanol were added to the nitrogen-saturated solution of
3-bromopyridine (2.66 g, 16.8 mmol) in toluene and refluxed under nitrogen for 8 hrs. Water was added to the solution and extracted with ethyl acetate, washed with water and brine and dried. Solvents were removed under reduced pressure and the residue was purified by silica gel column chromatography (eluent; hexane/ethyl acetate = 1/1.5) to give 4-(3-pyridyl)benzaldehyde (0.78 g, 25%).

Methyl iodide (0.8 ml, 12.9 mmol) was added to a solution of 4-(3-pyridyl)benzaldehyde (0.78 g, 4.3 mmol) in dichloromethane and stirred 2 days. Additional methyl iodide (0.8 ml, 12.9 mmol) was added and stirred for 3 hr. After removal of the solvent, methanol was added to the residue and ice-cooled. Sodium tetrahydroborate (6.4 g, 17.0 mmol) was added to the solution and stirred for 1.5 hr with warming to room temperature. Organic solvents were removed under reduced pressure after addition of water and extracted with ethyl acetate, washed with water and brine and dried over sodium sulfate. After removal of the solvent under reduced pressure, residue was purified by silica gel chromatography (eluent ethyl acetate to methanol) to give 3-(4-hydroxymethylphenyl)-1-methyl-1,2,5,6-tetrahydropyridine (0.63 g, 72%).

Triethylamine (1.29 ml, 9.2 mmol), acetic anhydride (0.35 ml, 3.7 mmol) were added to a solution of 4-(hydroxymethyl)phenyl-1-methyl-1,2,5,6-tetrahydropyridine (0.63 g, 3.1 mmol) in dichloromethane and stirred overnight.





A solution of 3-(4-acetoxymethylphenyl)-1-methyl-1,2,5,6tetrahydropyridine (0.67 g, 2.7 mmol) and 1-chloroethyl chloroformate (0.36 ml, 3.3
mmol) in dichloroethane was refluxed for 2 hr. Organic solvents were removed
under reduced pressure after addition of water and extracted with ethyl acetate,
washed with water and brine and dried over sodium sulfate. After removal of the
solvent, methanol was added and refluxed for 1.5 hr. Tetrahydrofuran and water
were added to the residue after removal of the solvent under reduced pressure and
triethylamine (1.9 ml, 13.6 mmol) and di-tert-butyl dicarbonate (0.66 g, 3.0 mmol)
were added and stirred. Organic solvents were removed under reduced pressure
after addition of water and extracted with ethyl acetate, washed with water and
brine and dried over sodium sulfate. Removal of the solvent under reduced pressure
and the residue was purified by silica gel chromatography to give
3-(4-acetoxymethylphenyl)-1-(tert-butoxycarbonyl)-1,2,5,6-tetrahydropyridine
(0.71 g, 78%).

Palladium on charcoal was added to the solution of 3-(4-acetoxy-methylphenyl)-1-(tert-butoxycarbonyl)-1,2,5,6-tetrahydropyridine (0.71 g, 2.1 mmol) in ethyl acetate and stirred under hydrogen atmosphere. After filtration with celite and removal of the solvent under reduced pressure, methanol and 1N aqueous sodium hydroxide were added and stirred. Organic solvents were removed under reduced pressure after addition of water and extracted with ethyl acetate, washed with water and brine and dried over sodium sulfate. Removal of the solvent under reduced pressure and the residue was purified by silica gel chromatography (eluent; hexane/ethyl acetate = 3/1) to give 3-(4-hydroxymethylphenyl)-1-(tert-butoxycarbonyl)piperidine (0.39 g, 62%).





Triethylamine (0.47 g, 3.4 mmol) and methanesulfonyl chloride (0.12 ml, 1.6 mmol) were added to an ice-cooled solution of 3-(4-hydroxymethylphenyl)-1-(tert-butoxycarbonyl)piperidine (0.39 g, 1.34 mmol) in dichloromethane and stirred for 7.5 hr. Pyrrolidine (1.0 ml, 12 mmol) was added to the solution and stirred overnight. Organic solvents were removed under reduced pressure after addition of water and extracted with ethyl acetate, washed with water and brine and dried over sodium sulfate. Removal of the solvent under reduced pressure and the residue was purified by silica gel chromatography (eluent; ethyl acetate to ethyl acetate/methanol = 1/1, then methanol only) to give 3-(4-(1-pyrrolidinyl)methyl-phenyl)-1-(tert-butoxycarbonyl)piperidine (0.26 g, 56%).

4N Hydrogen chloride in ethyl acetate was added to 3-(4-(1-pyrrolidinyl)-methylphenyl)-1-(tert-butoxycarbonyl)piperidine (0.26 g, 0.75 mmol) and stirred overnight. After filtration and dryness, triethylamine (0.5 ml, 3.6 mmol), 2-chloro-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (0.14 g, 0.63 mmol) and tetrahydrofuran were added and stirred at 70°C. Organic solvents were removed under reduced pressure after addition of water and extracted with ethyl acetate, washed with water and brine and dried over sodium sulfate. Removal of the solvent under reduced pressure and the residue was dissolved into ethyl acetate. A solution of fumaric acid (0.095 g, 0.82 mmol) in acetone was added and the resulting precipitate was filtered and dried to give the title compound (0.29 g, 76%).

Example 5: Synthesis of (R)-2-(2-(4-chlorophenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-pyrimidin-4- one (No. XA372)

To a solution of (S)-2-methyl-CBS-oxazaborolidine (27.6 mL, 1.0 M solution in toluene, 27.6 mmol) was added borane-tetrahydrofuran complex (166 ml, 1.0 M solution in tetrahydrofuran, 166 mmol) at -40 °C. To the resulting solution was added a solution of 4'-chlorophenacyl bromide (32.25 g, 138.1 mmol) in tetrahydrofuran (200 ml) through dropping funnel over 1 h at -40 °C. After stirring



4,144

WO 2004/085408

for 3 hours below 0 °C, methanol (ca. 50 ml) was added dropwise. After stirring the resulting solution for additional 30 min at room temperature, solvent was removed under reduced pressure. The residue, dissolved in ethyl acetate, was treated with 1 N hydrochloric acid to form white precipitate, which was filtered off. The layers of the filtrate was separated, and the organic layer was washed with hydrochloric acid and brine successively, dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure. The residue was used for the next reaction without further purification.

The residue was dissolved in ether (250 ml), and stirred with potassium hydroxide (15.5 g, 276 mmol) in water (250 ml) vigorously. After consumption of the starting material, the layers were separated. The organic layer was washed with brine, dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure. The residue was used for the next reaction without further purification.

The residue was heated with benzylamine (37.7 ml, 345 mmol) at 80 °C for 4.5 h. After cooling at room temperature, the resulting white crystals was washed with ether/hexane and collected to afford (S)-2-benzylamino-1-(4-chlorophenyl)-ethanol (23.8 g, 65.8%). The excess benzylamine in the filtrate was distilled off at 120 °C under reduced pressure. From the residue, another (S)-2-benzylamino-1-(4-chlorophenyl)ethanol (2.41 g, 6.7%) was obtained.

1H NMR (CDCl<sub>3</sub>) ™: 2.68(1H, dd, J=12.3, 8.9Hz), 2.92(1H, dd, J=12.3, 3.7Hz), 3.80(1H, d, J=11.9Hz), 3.86(1H, d, J=11.9Hz), 4.68(1H, dd, J=8.9, 3.7Hz), 7.30(9H, m).

To a suspension of (S)-2-benzylamino-1-(4-chlorophenyl)ethanol (15.76 g, 60.21 mmol) and triethylamine (33.6 ml, 241 mmol) in dichloromethane (300 ml) was added a solution of thionyl chloride (4.83 ml, 66.2 mmol) in dichloromethane (20 ml) at -78 °C over 20 min. The resulting suspension was stirred at -78 °C for 20 min and at 0 °C for additional 20 min. The reaction mixture was partitioned





between ether and water, and the organic layer was washed with brine, dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure. The residue was purified by silica gel column chromatography (eluent: 10-20% ethyl acetate-hexane) to afford (2RS,5S)-3-benzyl-5-(4-chlorophenyl)-1,2,3-oxathiazolidine 2-oxide (16.2 g 87.4%) as a pale yellow solid.

The resulting product was obtained as a mixture of two diastereomers due to the S-oxide.

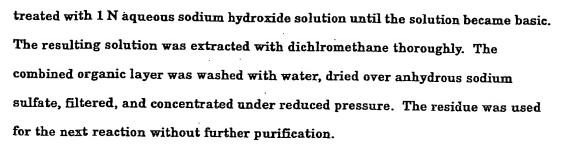
major isomer: <sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ : 3.31(1H, dd, J=10.5, 9.9Hz), 3.55(1H, dd, J=9.0, 6.3Hz), 3.88(1H, d, J=13.2Hz), 4.37(1H, d, J=13.2Hz), 5.49(1H, dd, J=10.5, 6.3Hz), 7.22-7.43(9H, m).

minor isomer: <sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ : 3.21(1H, dd, J=13.5, 4.5Hz), 3.77(1H, dd, J=13.5, 11.4Hz), 4.05(1H, d, J=13.5Hz), 4.38(1H, d, J=13.5Hz), 5.99(1H, dd, J=11.4, 4.5Hz), 7.22-7.43(9H, m).

A solution of (2RS,5S)-3-benzyl-5-(4-chlorophenyl)-1,2,3-oxathiazolidine 2-oxide (16.2 g, 52.6 mmol) and sodium azide (17.11 g, 263.2 mmol) in N,N-dimethylformamide (100 ml) was heated at 70 °C for 24 hours. The reaction mixture was partitioned between ether and water, and the organic layer was washed with water and brine successively, dried over anhydrous sodium sulfate, and concentrated under reduced pressure. The residue was purified by silica gel column chromatography (eluent: 10-20% ethyl acetate-hexane) to afford (R)-N-benzyl-2-azido-2-(4-chlorophenyl)ethylamine (12.7 g, 83.8%) as a yellow oil. 1H NMR (CDCl<sub>3</sub>) δ: 2.81(1H, dd, J=12.5, 5.1Hz), 2.89(1H, dd, J=12.5, 8.5Hz), 3.82(2H, s),4.64(1H, dd, J=8.5, 5.1Hz),7.23-7.36(9H, m).

A solution of (R)-N-benzyl-2-azido-2-(4-chlorophenyl)ethylamine (12.7 g, 44.1 mmol) in tetrahydrofuran (176 mL) was treated with triphenylphosphine (13.9 g, 52.9 mmol) at room temperature. After addition of water (20 ml), the reaction mixture was heated at 60 °C for 1 h. The reaction mixture was condensed, and partitioned between ether and 1 N hydrochloric acid. The aqueous layer was





The residue was heated with diethyl oxalate (18 ml, 132 mmol) at 120 °C for 1.5 h. The resulting white precipitate was washed with ether and collected to afford (R)-1-benzyl-5-(4-chlorophenyl)-2,3-dioxopiperazine (11.4 g, 82.2%). 

<sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ : 3.46(1H, dd, J=12.9, 8.1Hz), 3.60(1H, dd, J=12.9, 3.8Hz), 4.48(1H, d, J=14.7Hz), 4.79(1H, d, J=14.7Hz), 4.80(1H, dd, J=8.9, 3.8Hz), 6.83(1H, s), 7.13(4H, m), 7.27(5H, m).

To a suspension of (R)-1-benzyl-5-(4-chlorophenyl)-2,3-dioxopiperazine (11.4 g, 36.3 mmol) in tetrahydrofuran (300 ml) was added borane-tetrahydrofuran complex (181 mL, 1.0 M solution in tetrahydrofuran, 181 mmol) at room temperature. After stirring for 24 hours, the reaction mixture was quenched with methanol (50 ml) at 0 °C, and concentrated under reduced pressure. The residue was treated with 10% aqueous sodium hydroxide solution (300 ml) and heated at 100 °C for 2 hours. After cooling at room temperature, the mixture was extracted with dichloromethane thoroughly. The combined organic layer was dried over anhydrous sodium sulfated, filtered, and concentrated under reduced pressure. The residue was used for the next reaction without further purification.

To a solution of the residue and triethylamine (7.58 ml, 54.4 mmol) in dichloromethane (150 ml) was added di-tert-butyl dicarbonate (9.49 g, 43.5 mmol) at room temperature. After stirring for 45 min, the resulting mixture was partitioned between dichloromethane and water, dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure. The residue was purified by silica gel column chromatography (eluent: 10-20% ethyl acetate-hexane) to afford (R)-1-benzyl-4-(tert-butoxycarbonyl)-3-(4-chlorophenyl)piperazine (11.6 g,





82.8%) as an oil.

WO 2004/085408

<sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ : 1.43(9H, s), 2.16(1H, dt, J=4.4, 11.7Hz), 2.40(1H, dd, J=4.4, 11.7Hz), 2.78(1H, dd, J=4.4, 11.7Hz), 2.98(1H, dt, J=4.4, 11.7Hz), 3.20(1H, d, J=12.8Hz), 3.42(1H, d, J=12.9Hz), 3.57(1H, d, J=12.9Hz), 3.89(1H, d, J=12.8Hz), 5.17(1H, s), 7.24-7.36(9H, m).

To a solution of (R)-1-benzyl-4-(tert-butoxycarbonyl)-3-(4chlorophenyl)piperazine (11.6 g, 30.1 mmol) in 1,2-dichloroethane (80 ml) was added 1-chloroethyl chloroformate (4.91 ml, 45.1 mmol) at room temperature. Upon disappearance of the starting material, the reaction mixture was concentrated under reduced pressure. The residue was then dissolved in methanol (100 ml) and refluxed for 30 min. The resulting white precipitate was filtered and washed with methanol to afford (R)-2-(4-chlorophenyl)piperazine dihydrochloride, which was liberated with aqueous sodium hydroxide solution, and extracted with dichloromethane to afford (R)-2-(4-chlorophenyl)piperazine (3.04 g, 51.4%) as white solid.

<sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ :2.65(1H, dd, J=12.0, 10.5Hz), 2.82-3.04(4H, m), 3.09(1H, d, J=12.6Hz), 3.73(1H, dd, J=10.1, 2.7Hz), 7.29(4H, m)

The filtrate was concentrated under reduced pressure and partitioned between ether and 1 N hydrochloric acid. The aqueous layer was neutralized with 1 N aqueous sodium hydroxide solution, and extracted with dichloromethane thoroughly. The combined organic extracts were dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure. The residue was purified after Boc-protection (Boc2O, Et3N, CH2Cl2) to furnish (R)-1,4-di(tert-butoxycarbonyl)-2-(4-chlorophenyl)piperazine (2.70 g, 22.6%) as pale yellow solid.

<sup>1</sup>H NMR (CDCl<sub>3</sub>)  $\delta$ : 1.43(9H, s), 1.46(9H, s), 2.96(2H, m), 3.32(1H, dd, J=13.8, 4.2Hz), 3.74(1H, m), 3.94(1H, d, J=11.4Hz), 4.40(1H, d, J=13.2Hz), 5.23(1H, s),7.25(2H, m)



To a suspension of (R)-2-(4-chlorophenyl) piperazine dihydrochloride (1.09 g, 4.05 mmol) in tetrahydrofuran (24 ml) was added triethylamine (2.82 ml, 20.3 mmol). After stirring for 15 min at room temperature, 2-chloro-3-methyl-6-(4pyridyl)-3H-pyrimidin-4-one (748 mg, 3.38 mmol) was added portionwise. Upon disappearance of the chloropyrimidone, the reaction mixture was condensed under reduced pressure. The residue was partitioned between saturated aqueous sodium bicarbonate solution and dichloromethane. The organic layer was dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure to give pale yellow solid, which was recrystallized from ethanol to afford (R)-2-(2-(4-chlorophenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (998 mg, 77.4%) as white crystals. The enantiomer excess was determined by HPLC (>99% ee). The crystals were converted into its dihydrochloride salt. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>)  $\delta$ : 3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) MS: 382(M+H)  $[\alpha]_{D^{24}} = +62.2 \circ (c\ 1.00,\ H_2O)$ 

Example 6: Synthesis of (S)-2-(2-(4-chlorophenyl)piperazin-4-yl)-3-methyl-6-(4-pyridyl)-pyrimidin-4-one (No. XA373)

(S)-isomer was prepared same as above by using (R)-2-methyl-CBS-oxazaborolidine instead of (S)-2-methyl-CBS-oxazaborolidine. 

<sup>1</sup>H NMR (DMSO-d<sub>6</sub>)  $\delta$ : 3.40 (3H, m), 3.45 (3H, s), 3.53-3.96 (3H, m), 4.68 (1H, t, J = 13.5Hz), 7.10 (1H, s), 7.60 (2H, d, J=8.3Hz), 7.76 (2H, d, J=8.3Hz), 8.38 (1H, br s), 8.91 (1H, d, J=4.8Hz), 9.88 (1H, br s), 10.31 (1H, br s)

MS: 382(M+H)

[ $\alpha$ ] $\alpha$ <sup>24</sup> = -63.3 ° (c 1.00, H<sub>2</sub>O)

Example 7: Synthesis of 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyrimidyl)-pyrimidin-4-one (No. YA0366)

A solution of 2-bromo-5-fluoroanisole (11.8 g, 57.6 mmol) in tetrahydrofuran (60 ml) was dropped into the magnesium (1.40 g, 57.6 mmol) in refluxed tetrahydrofuran (32 ml) containing small amount of 1,2-dibromoethane and refluxed for 15 min. After addition of tetrahydrofuran (50 ml), the solution was cooled to -78 °C and diethyl oxalate (7.41 g, 50.7 mmol) in diethyl ether (50 ml) was dropped into the solution. After stirring at the same temperature for 30 min, the solution was warmed to -10°C and 1N aqueous hydrogen chloride (50 ml) and water were added. Organic layer was extracted with diethyl ether, washed with brine and dried over magnesium sulfate. After removal of the solvent under reduced pressure, purification of the residue by silica gel column chromatography (eluent: hexane/ethyl acetate = 5/2) gave ethyl 2-(4-fluoro-2-methoxyphenyl)-2-oxoacetate (6.80g, 59%)

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 1.40(3H, t, J=7.1 Hz),3.87(3H, s), 4.89(2H, q, J=7.1Hz), 6.68(1H, d, J=10.5 Hz), 6.77-6.81(1H, m), 7.91-7.95(1H, m).

Ethylenediamine (0.60 g, 10.0 mmol) was added to a solution of ethyl 2-(4-fluoro-2-methoxyphenyl)-2-oxoacetate (2.26 g, 10.0 mmol) in ethanol(30 ml) and refluxed 4 hr. After removal of the solvent under reduced pressure, residue was washed with ethanol-diethyl ether to give 5,6-dihydro-3-(4-fluoro-2-methoxyphenyl)pyrazinone (1.76 g, 79%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 3.50-3.56 (2H, m), 3.81 (3H, s), 3.88-3.92 (2H, m), 6.65(1H, d, J=11.0 Hz), 6.70-6.76 (1H, m), 6.89(1H, bs), 7.36-7.40(1H, m).

5,6-Dihydro-3-(4-fluoro-2-methoxyphenyl)pyrazinone was added to the solution of lithium aluminium hydride (0.46 g, 12 mmol) in diethyl ether (25 ml) and refluxed for 6 hr. Water (0.48 ml), 15% sodium hydroxide solution (0.48 ml) and again water (1.21 ml) were added to the ice-cooled solution and the precipitate was

filtered and washed with dichloromethane. Combined organic layer was evaporated to give 2-(4-fluoro-2-methoxyphenyl)piperazine (0.83 g, 99%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 2.02(2H, s), 2.57-2.63 (1H, m), 2.80-2.89 (1H, m), 2.92-2.99 (2H, m), 3.06-3.12 (2H, m), 3.80(3H, s), 4.06 (1H, d, J=10.0 Hz), 6.56-6.65 (2H, m), 7.40 (1H, t, J=7.8 Hz).

2-Chloro-3-methyl-6-(4-pyrimidyl)-pyrimidin-4-one (223 mg, 1:0 mmol) was added to an ice-cooled solution of 2-(4-fluoro-2-methoxyphenyl)piperazine (210 mg, 1.0 mmol), triethylamine (0.15 ml, 1.1 mmol) in N,N-dimethylformamide (10 ml) and stirred at that temperature for 0.5 hr and then at room temperature for 3 hours. Reaction was quenched by ice-water and the filtrate was washed with water and dried to give 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyrimidyl)-pyrimidin-4-one (262 mg, 66%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 2.89-2.98 (1H, m), 3.22-3.31 (3H, m), 3.60 (3H, s), 3.62-3.71 (2H, m), 3.86 (3H, s), 4.39-4.44 (1H, m), 6.43-6.73 (2H, m), 7.33 (1H, s), 7.52-7.56 (1H, m), 8.19 (1H, d, J=5.1 Hz), 8.87 (1H, d, J=5.2 Hz), 9.28 (1H, d, J=1.2 Hz).

4N Hydrogen chloride in 1,4-dioxane (0.2 ml) was added to the solution of 2-(2-(4-fluoro-2-methoxyphenyl)piperazin-4-yl)-3-methyl-6-(4-pyrimidyl)-pyrimidi n-4-one (238 mg, 0.6 mmol) in dichloromethane (5 ml) and stirred for 15 min. Wash with methanol and ethyl acetate after removal of the solvent and dryness gave the title compound (223 mg, 86%).

Example 8: Synthesis of 2-(2-(4-chlorophenyl)-piperazine-4-yl)-3-methyl-6-(4-pyrimidyl)pyrimidin-4-one (No. YA0269)

Dimethyl sulfoxide (60 ml) solution of 4-chlorophenacylbromide (11.11 g, 65.9 mmol) and water (1.7 ml) were stirred. The solution was extracted with ethyl acetate 3 times and washed with water twice and brine and then dried over sodium sulfate. After removal of the solvent, the residue was washed with hexane-ethyl acetate and dried to give 4-chlorophenylglyoxal (4.43 g, 50%).

<sup>1</sup>H-NMR (CDCl<sub>3</sub>)  $\delta$ : 4.02-4.16(2H, m), 5.90-5.95(1H, m), 7.45-7.53(2H, m), 8.05-8.11(2H, m).

A methanol (10 ml) solution of ethylenediamine (1.90 g, 31.6 mmol) was added to the ice-cooled solution of 4-chlorophenylglyoxal (4.43 g, 26.3 mmol) in methanol (100 ml) and tetrahydrofuran (30 ml) and stirred for 10 min. After addition of sodium tetrahydroborate (3.26 g, 86.3 mmol), additional methanol (50 ml) was added and stirred overnight. After removal of the solvent, diluted hydrochloric acid was added and extracted with ether twice. After addition of sodium hydroxide, basic aqueous layer was extracted with dichloromethane three times and washed with brine and dried over sodium sulfate. After removal of the solvent by filtration, purification of the residue by silica gel column chromatography (eluent; dichloromethane/ethanol = 10/1 to dichloromethane/ethanol/diethylamine = 20/2/1) to give 2-(4-chlorophenyl)-piperazine (0.43 g, 9%)

1H-NMR (CDCl<sub>3</sub>)  $\delta$ : 2.67(1H, dd, J=10.5, 12.0 Hz), 2.87-3.03(4H, m), 3.07-3.13(1H, m), 3.77(1H, dd, J=2.7, 10.2 Hz), 7.27-7.36(4H, m).

Triethylamine (528 mg, 5.2 mmol) was added to a solution of 4-(chlorophenyl)piperazine (216 mg, 1.1 mmol) and 2-chloro-3-methyl-6-(4-pyrimidyl)pyrimidin-4-one and stirred at 50°C for 2 hr. Solvent was removed under reduced pressure, and 1N aqueous sodium hydroxide solution was added to the residue and extracted by dichloromethane. After washing with brine and dryness by sodium sulfate, solvent was removed under reduced pressure, and the residue was purified using ISOLUTE(registered trade mark) SI (International Sorvent Technology, UK)(eluent; dichloromethane/ethanol = 10/1) to give the title compound (396 mg, 95 %).

Example 9: Synthesis of 2-(2-(4-chlorophenyl)-6,6-dimethyl-piperazin-4-yl)-3-methyl-6-pyridin-4-yl-3*H*-pyrimidin-4-one dihydrochloride (No. XA1986)

A solution of 4'-chloro-2-bromoacetophenone (25.0 g, 107 mmol), water (1.92 mL, 107 mmol) and 47% hydrobromic acid (0.20 mL) in dimethylsulfoxide (160 mL) was stirred at 80°C for 5 h. After the reaction mixture was poured into water, the precipitate was filtered, washed with diethylether and dried, affording 4'-chloro-2,2-dihydroxyacetophenone (14.0 g, 70%). <sup>1</sup>H NMR (300MHz, CDCl<sub>3</sub>), δ 5.92(1H, s), 7.45-7.52(2H, m), 8.05 –8.20(2H, m).

2,2-dimethly-ethylenediamine (2.10 mL, 20.0 mmol) was added to a solution of 4'-chloro-2,2-dihydroxyacetophenone (3.70 g, 20.0 mmol) in methanol (120 mL) and tetrahydrofuran (30 mL) at room temperature. After 2 h, sodium borohydride (1.50 g, 40.0 mmol) was added to the reaction mixture at 0  $^{\circ}$ C. The reaction mixture was stirred overnight, then quenched with 1N hydrochloric acid and evaporated in vacuo. The acidic solution was extracted with ethyl acetate, then basified to pH 11 using 15% aqueous sodium hydroxide, and extracted with dichloromethane. The extract was dried over sodium sulfate and concentrated in vacuo. Di-t-butyldicarbonate (6.40 mL, 27.9 mmol) was added to the solution of the residue in 1N aqueous sodium hydroxide (40 mL) and tetrahydrofuran (60 mL). The resulting suspension was heated at 40  $^{\circ}$ C for 8 h, then diluted with ethyl acetate and water. The organic layer was extracted with additional ethyl acetate, dried and concentrated in vacuo. The crude product was purified by flash column chromatography, affording 2-(4-chlorophenyl)-4-t-butoxycarbonyl-6,6-dimethylpiperazine (1.69 g, 28%, 2 steps).  $^{1}$ H NMR (300MHz, CDCl<sub>3</sub>),  $\delta$  1.15(3H, s), 1.21(3H, s), 2.47-2.70(2H, m), 3.72-4.16(3H, m), 7.26-7.37(4H, m).

4 M Hydrogen chloride in ethyl acetate (5.0 mL, 20.0 mmol) was added to a solution of 2-(4-chlorophenyl)-4-t-butoxycarbonyl-6,6-dimethyl-piperazine (1.69 g, 5.2 mmol). After 12 h, removing the solvent, filtrating and washing the precipitate with ethyl acetate gave 2-(4-chlorophenyl)-6,6-dimethyl-piperazine dihydrochloride

(1.43 g, 95%). <sup>1</sup>H·NMR (300MHz, DMSO-d<sub>6</sub>),  $\delta$  1.40 (3H, s), 1.58(3H, s), 3.24-3.99(4H, m), 4.73(1H, m), 7.69(2H, d, J = 8.4 Hz), 7.79(2H, m), 9.99-10.12(2H, m).

A solution of 2-(4-chlorophenyl)-6,6-dimethyl-piperazine hydrochloride (155 mg, 0.52 mmol), 2-chloro-3-methyl-6-(4-pyridyl)-pyrimidine-4-one (111 mg, 0.50 mmol) and triethylamine (0.42 mL, 2.50 mmol) in tetrahydrofuran (5 mL) was stirred at room temperature for 6 h. The whole was evaporated in vacuo and the residue was extracted with dichloromethane. The organic layer was washed with water, dried and concentrated in vacuo. The residue was dissolved in methanol (5mL) and treated with 4M hydrogen chloride in ethyl acetate (0.50 mL, 2.0 mmol) for 20 min. After removing the solvent, filtrating and washing the precipitate with ethanol gave 2-(2-(4-chlorophenyl)-6,6-dimethyl-piperazin-4-yl)-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one dihydrochloride (235 mg, 97%).

Example 10: Synthesis of 2-(2S-(4-bromophenyl)-piperazin-1-yl)-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one (No. XA2051)

Benzyl chloroformate (2.40 mL, 15.0 mmol) was added to a solution of 2S-(4-bromophenyl)-piperazine dihydrochloride in 1N aqueous sodium hydroxide (30 mL) and dichloromethane (60 mL). The resulting suspension was stirred at room temperature for 1.5 h. After partitioned between ethyl acetate, the organic layer was extracted with additional ethyl acetate, dried and concentrated in vacuo. The precipitate was washed with ether, affording 2S-(4-bromophenyl)-4-benzyloxycarbonyl-piperazine (2.92 g, 57%). <sup>1</sup>H NMR (300MHz, CDCl<sub>3</sub>), δ 2.87-3.01(2H, m), 3.47(2H, m), 3.93-3.97(1H, m), 4.20(2H, m), 5.16(2H, s), 7.36(5H, m), 7.42-7.61(4H, m).

A solution of 2S-(4-bromophenyl)-4-benzyloxycarbonyl-piperazine (788 mg, 2.10 mmol), 2-chloro-3-methyl-6-(4-pyridyl)-pyrimidine-4-one (444 mg, 2.00 mmol) and diisopropylethylamine (0.70 mL, 4.00 mmol) in dimethylformamide (20 mL) was stirred at 80°C for 3 h. The reaction mixture was poured into water and the

whole was extracted with ethyl acetate. The organic layer was washed with brine, dried and concentrated in vacuo. Chromatographic purification of the residue provided 2-(2S-(4-bromophenyl)-4-benzyloxycarbonyl-piperazin-1-yl)}-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one (601 mg, 54%). <sup>1</sup>H NMR (300MHz, CDCl<sub>3</sub>),  $\delta$  3.05(1H, m), 3.30-3.48(3H, m), 3.64(3H, s), 4.08-4.22(2H, m), 4.68(1H, m), 5.15(1H, d, J= 12.3 Hz), 5.21(1H, d, J= 12.6 Hz), 6.63(1H, s), 7.21(2H, d, J= 8.4 Hz), 7.28-7.39(7H, m), 7.59(2H, d, J=6.3 Hz), 8.68(2H, d, J=6.3 Hz).

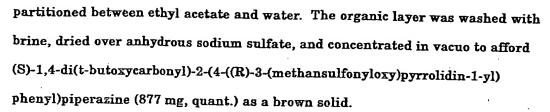
Potassium hydroxide (168 mg, 3.0 mmol) was added to a solution of 2-{2S-(4-bromophenyl)-4-benzyloxycarbonyl-piperazin-1-yl}-3-methyl-6-pyridin-4-y l-3H-pyrimidin-4-one in ethanol (2.0 mL). After stirring for 8 h at room temperature, purifying by preparative HPLC gave 2-(2S-(4-bromophenyl)-piperazin-1-yl)-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one (40 mg, 26%).

Example 11: Synthesis of (S)-3-methyl-6-(4-pyridyl)-2-(3-(4-(3-(pyrrolidin-1-yl) pyrrolidin-1-yl)phenyl)piperazin-1-yl)pyrimidin-4-one (No. XA2032)

A suspension of (S)-2-(4-bromophenyl)-1,4-di(t-butoxycarbonyl) piperazine (1.33 g, 3.00 mmol), (R)-3-pyrrolidinol (520 mg, 4.20 mmol), palladium acetate (27 mg, 0.12 mmol), 2-(di-t-butylphosphino)biphenyl (72 mg, 0.24 mmol), and sodium t-butoxide (808 mg, 8.41 mmol) in tert-butanol (20 mL) was heated at 90 °C for 3.5 h. After dilution with ethyl acetate, the resulting mixture was passed through a Celite column. The filtrate was concentrated in vacuo, and the residue was purified by silica gel column chromatography eluting 10-50% ethyl acetate - hexane to afford (S)-1,4-di-(t-butoxycarbonyl)-2-(4-((R)-3-hydroxypyrrolidino) phenyl)piperazine (733 mg, 54.5%) as a yellow foam.

To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4-((R)-3-hydroxy pyrrolidino) phenyl)piperazine (733 mg, 1.64 mmol) and triethylamine (0.34 mL, 2.46 mmol) in dichloromethane (20 mL) was added methanesulfonyl chloride (0.152 mL, 1.97 mmol) at 0 °C. After stirring for 20 min, the reaction mixture was

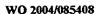




To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4-((R)-3-methansulfonyloxy-pyrrolidino)phenyl)piperazine (877 mg, 1.64 mmol) in toluene (10 mL) was added pyrrolidine (0.64 mL, 8.19 mmol), and the resulting solution was heated at 90 °C for 8 h. After checking consumption of the starting material with TLC, the reaction mixture was partitioned between ethyl acetate and saturated sodium bicarbonate aqueous solution. The organic layer was washed with brine, dried over anhydrous sodium sulfate, and concentrated in vacuo. The residue was purified by silica gel column chromatography eluting 30-100% ethyl acetate-hexane and then 3-10% methanol-ethyl acetate to afford (S)-1,4-di(t-butoxycarbonyl)-2-(4-((S)-3-(pyrrolidin-1-yl)pyrrolidin-1-yl) phenyl)piperazine (479 mg, 58%) as a pale yellow powder.

To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4-((S)-3-(pyrrolidin-1-yl) pyrrolidin-1-yl)phenyl)piperazine (479 mg, 0.957 mmol) in dichloromethane (4 mL) was added 4 N hydrogen chloride in ethyl acetate (4 mL) at room temperature. After stirring for 3 h, the resulting precipitate was collected and dried in vacuo to afford (S)-2-(4-((S)-3-(pyrrolidin-1-yl)pyrrolidin-1-yl)phenyl)piperazine tetrahydrochloride (370 mg, 87%) as a white solid.

To a suspension of (S)-2-(4-((S)-3-(pyrrolidin-1-yl)pyrrolidin-1-yl)phenyl) piperazine tetrahydrochloride (98 mg, 0.22 mmol) in tetrahydrofuran (5 mL) was added triethylamine (0.20 mL, 1.40 mmol) and 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (44 mg, 0.20 mmol) at room temperature. After stirring for 24 h, the reaction mixture was concentrated in vacuo. The residue was dissolved in dichloromethane and sodium bicarbonate aqueous solution, and the solution was passed through CHEM ELUT CE1010 (manufactured by VARIAN). The filtrate was



concentrated, and the resulting crystals were washed in a mixture of diisopropyl ether and ethanol to afford (S)-2-(3-(4-(3-(pyrrolidin-1-yl)pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3- methyl-6-(4-pyridyl)pyrimidin-4-one (80 mg, 82%) as a pale yellow solid.

Example 12: Synthesis of (S)-3-methyl-6-(4-pyrimidinyl)-2-(3-(4-(3-(pyrrolidin-1-yl) pyrrolidin-1-yl)phenyl)piperazin-1-yl)pyrimidin-4-one (No. YA1577)

To a suspension of (S)-2-(4-((S)-3-(pyrrolidin-1-yl)pyrrolidin-1-yl)phenyl) piperazine tetrahydrochloride (99 mg, 0.22 mmol) in tetrahydrofuran (5 mL) was added triethylamine (0.20 mL, 1.40 mmol) and 2-chloro-3-methyl-6-(4-pyrimidinyl)-3H-pyrimidin-4-one (45 mg, 0.20 mmol) at room temperature. After stirring for 24 h, the reaction mixture was concentrated in vacuo. The residue was dissolved in dichloromethane and sodium bicarbonate aqueous solution, and the solution was passed through CHEM ELUT CE1010 (manufactured by VARIAN). The filtrate was concentrated, and the resulting crystals were washed in a mixture of diisopropyl ether and ethanol to afford (S)-3-methyl-6-(4-pyrimidinyl)-2-(3-(4-(3-(pyrrolidin-1-yl)pyrrolidin-1-yl)phenyl)piperazin-1-yl)-pyrimidin-4-one (65 mg, 66%) as a pale yellow solid.

Example 13: Synthesis of (S)-2-(3-(4-(N-cyclohexyl-N-methylamino)phenyl) piperazin- 1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (No. XA1999)

A suspension of (S)-2-(4-bromophenyl)-1,4-di(t-butoxycarbonyl) piperazine (1.21 g, 2.75 mmol), N-methylcyclohexylamine (0.43 mL, 3.30 mmol), palladium acetate(25 mg, 0.11 mmol), 2-(di-t-butylphosphino)biphenyl (66 mg, 0.22 mmol), and sodium t-butoxide (370 mg, 3.85 mmol) in t-butanol (15 mL) was heated at 80 °C for 8 h. The resulting solution was partitioned between ethyl acetate and water. The organic layer was washed with brine, dried over anhydrous sodium sulfate, and concentrated in vacuo. The residue was purified by silica gel column





chromatography eluting 10-15% ethyl acetate-hexane to afford (S)-1,4-di(t-butoxycarbonyl)-2-(4-(N-cyclohexyl-N-methylamino)phenyl)piperazine (917 mg) as white crystals.

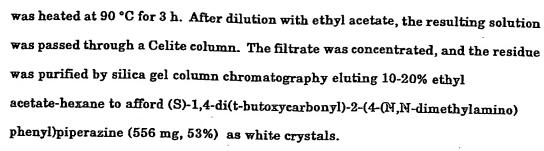
To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4-(N-cyclohexyl-N-methylamino)phenyl)piperazine in dichloromethane (4 mL) was added 4 N hydrogen chloride in ethyl acetate(4 mL). After stirring for 40 min, the white precipitate was collected, which included impurities. The mixture was purified by a reverse phase chromatography eluting 0.05% TFA in water-acetonitrile to afford (S)-2-(4-(N-cyclohexyl-N-methylamino)phenyl)piperazine (59 mg 8% in 2 steps) as a clear oil.

To a solution of (S)-2-(4-(N-cyclohexyl-N-methylamino)phenyl) piperazine(50 mg, 0.183 mmol) and triethylamine (0.077 mL, 0.55 mmol) was added 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (37 mg, 0.17 mmol) at room temperature. After stirring for 4.5 h, the reaction mixture was concentrated in vacuo. The residue was partitioned between dichloromethane and saturated sodium bicarbonate aqueous solution. The organic layer was dried over anhydrous sodium sulfate and concentrated. The residue was purified by a reverse phase chromatography eluting 0.05% TFA in water-acetonitrile to afford (S)-2-(3-(4-(N-cyclohexyl-N-methylamino)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyr idyl)pyrimidin-4-one (67 mg, 88%) as a oil, which was dissolved in ethyl acetate and treated with 4 N hydrogen chloride in ethyl acetate to yield its trihydrochloride.

Example 14: Synthesis of (S)-2-(3-(4-(N,N-dimethylamino)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one trihydrochloride (No. XA2017)

A suspension of (S)-2-(4-bromophenyl)-1,4-di(t-butoxycarbonyl) piperazine (1.14 g, 2.59 mmol), N,N-dimethylamine hydrochloride (422 mg, 5.17 mmol), palladium acetate (23 mg, 0.10 mmol), 2-(di-t-butylphosphino)biphenyl(62 mg, 0.21 mmol), and sodium t-butoxide (845 mg, 8.80 mmol) in t-butanol (15 mL)





To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4-(N,N-dimethylamino) phenyl)piperazine (556 mg, 1.37 mmol) in dichloromethane (4 mL) was added 4 N hydrogen chloride in ethyl acetate (4 mL). After stirring for 8.5 h, the white precipitate was collected and dried in vacuo to afford (S)-2-(4-(N,N-dimethylamino) phenyl)piperazine trihydrochloride (413 mg, 96%) as white crystals.

To a suspension of (S)-2-(4-(N,N-dimethylamino)phenyl)piperazine trihydrochloride(115 mg, 0.365 mmol) in tetrahydrofuran (5 mL) was added triethylamine (0.28 mL, 2.0 mmol) and then 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (74 mg, 0.33 mmol) at room temperature. After stirring for 10 h, the resulting mixture was concentrated in vacuo. The residue was dissolved in dichloromethane and saturated sodium bicarbonate aqueous solution, and the solution was passed through CHEM ELUT CE1010 (manufactured by VARIAN). The filtrate was concentrated in vacuo to yield crystals, which were washed with diisopropyl ether. After the crystals were dissolved in ethyl acetate, the solution was treated with 4 N hydrogen chloride in ethyl acetate. White precipitate was collected and dried in vacuo to afford (S)-2-(3-(4-(N,N-dimethylamino)phenyl) piperazin-1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one trihydrochloride (135 mg, 81%).

Example 15: Synthesis of (S)-2-(3-(4-methoxybiphen-4-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (No. XA1991)

A mixture of (S)-2-(4-bromophenyl)-1,4-di(t-butoxycarbonyl)piperazine (1.82 g, 4.11 mmol), 4-methoxyphenylboronic acid (937 mg, 6.17 mmol), sodium



carbonate (2.18 g, 20.6 mmol), and tetrakis(triphenylphosphine)palladium(0) (238 mg, 0.206 mmol) was dissolved in dimethoxyethane (20 mL) and water (20 mL), and the resulting solution was refluxed for 3 h. After cooling to room temperature, the mixture was partitioned between ethyl acetate and water. The organic layer was washed with brine, dried over anhydrous sodium sulfate, and concentrated in vacuo. The resulting solid was washed with ethyl acetate to afford (S)-1,4-di(t-butoxycarbonyl)-2-(4'-methoxybiphen-4-yl) piperazine (1.46 g, 75.9%) as a white solid.

To a solution of (S)-1,4-di(t-butoxycarbonyl)-2-(4'-methoxybiphen-4-yl)-piperazine (1.46 g, 3.12 mmol) in dichloromethane (8 mL) was added 4 N hydrogen chloride in ethyl acetate (8 mL) at room temperature. After stirring for 1 h, the precipitate was collected and dried in vacuo to afford (S)-2-(4'-methoxybiphen-4-yl) piperazine dihydrochloride (1.00 g, 94%) as white solid.

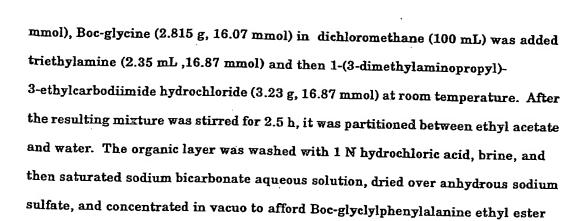
To a suspension of (S)-2-(4'-methoxybiphen-4-yl)-piperazine dihydrochloride (237 mg, 0.694 mmol) in tetrahydrofuran (5 mL) was added triethylamine (0.40 mL, 2.9 mmol) and then 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (128 mg, 0.579 mmol) at room temperature. After stirring for 28 h, the resulting mixture was concentrated in vacuo. The residue was partitioned between dichloromethane and saturated sodium bicarbonate aqueous solution, and the organic layer was dried over anhydrous sodium sulfate and then concentrated in vacuo. The resulting solid was washed with hot ethanol to afford (S)-2-(3-(4-methoxybiphen-4-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (252 mg, 96%), which was treated with 4 N hydrogen chloride in ethyl acetate to yield its dihydrochloride salt (252 mg) as pale yellow crystals.

Example 16: Synthesis of (S)-2-(3-benzylpiperazin-1-yl)-3-methyl-6-(4-pyridyl) pyrimidin-4-one (No. XA2004)

To a solution of L-phenylalanine ethyl ester hydrochloride (3.875 g, 16.87



(5.96 g).



To a solution of Boc-glycylphenylalanine ethyl ester (5.96 g) in dichloromethane (20 ml) was added trifluoroacetic acid (20 mL) at room temperature. After stirring 1.5 h, the resulting solution was concentrated in vacuo. The residue was dissolved in water, into which sodium bicarbonate was added until the pH was 9. After the solution was stirred for several hours, the resulting white crystals were collected and dried in vacuo to afford (S)-3-benzyl-2,5-dioxopiperazine (2.29 g, 70% in 2 steps) as a white powder.

To a suspension of (S)-3-benzyl-2,5-dioxopiperazine (2.284 g, 11.18 mmol) in tetrahydrofuran (20 mL) was added borane-tetrahydrofuran complex (49 mL, 1.0 M solution in THF, 49 mmol) at room temperature. The resulting mixture was refluxed for several hours before it was quenched with methanol at 0 °C. After concentration in vacuo, the residue was treated with 10% sodium hydroxide aqueous solution, which was extracted with dichloromethane thoroughly. The organic layer was washed with water, dried over anhydrous sodium sulfate, and concentrated in vacuo to afford white crystals, which were washed with ether to yield (S)-2-benzylpiperazine (795 mg, 40.3%).

To a solution of (S)-2-benzylpiperazine (48 mg, 0.27 mmol) in tetrahydrofuran (5 mL) was added triethylamine (0.10 mL, 0.74 mmol) and then 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (55 mg, 0.248 mmol) at room temperature. After refluxing for 24 h, the resulting mixture was concentrated in



vacuo. The residue was partitioned between dichloromethane and saturated sodium bicarbonate aqueous solution, and the organic layer was dried over anhydrous sodium sulfate and then concentrated in vacuo. The residue was purified by a reverse phase chromatography eluting 0.05% TFA in water-acetonitrile to afford (S)-2-(3-benzylpiperazin-1-yl)-3-methyl-6-(4-pyridyl) pyrimidin-4-one (73 mg 81%), which was treated with 4 N hydrogen chloride in ethyl acetate to yield its dihydrochloride salt as a yellow powder.

Example 17: Synthesis of (S)-3-methyl-2-(3-(4-(1,2,4-oxadiazol-3-yl)phenyl) piperazin-1-yl)-6-(4-pyridyl)pyrimidin-4-one (No. XA2039)

To a solution of 4-cyanoacetophenone (11.32 g, 77.98 mmol) in dichloromethane (200 mL) was added bromine (4.00 mL, 78.0 mmol) dropwise at room temperature. After stirring several minutes, the reaction mixture was washed with water, dried over anhydrous sodium sulfate, and concentrated in vacuo to afford 4-cyanophenacyl bromide (17.73 g) as a white solid.

A solution of 4-cyanophenacyl bromide (11.20 g, 49.99 mmol) in dimethylsulfoxide (83 mL) was treated with water (0.90 mL, 49.99 mmol). After stirring for 24 h at room temperature, it was poured into ice-water, and extracted with ether. The organic layer was washed with water and then brine, dried over anhydrous sodium sulfate, and concentrated in vacuo. The residue was purified by a silica gel column chromatography eluting 20-50% ethyl acetate in hexane to afford 4-cyanophenylglyoxal (5.10 g, 64.1%) as a yellow solid.

To a solution of 4-cyanophenylglyoxal (2.21 g, 12.5 mmol) in methanol (30 mL) and tetrahydrofuran (10 mL) was added ethylenediamine (1.00 mL, 14.96 mmol) at room temperature. After the mixture was stirred at room temperature for 1 h, sodium borohydride (943 mg, 24.92 mmol) was added at 0 °C. The solution was warmed up to room temperature and stirred for another 2 h before it was quenched with 1 N hydrochloric acid. After concentration in vacuo, the mixture was

partitioned between ether and water. The aqueous layer was alkalized with sodium hydroxide, and extracted with dichloromethane. The extract was dried over anhydrous sodium sulfate, and then concentrated in vacuo to afford reddish oil (1.69 g). The oil was dissolved in dichloromethane (30 mL), into which triethylamine (3.82 mL, 27.41 mmol) and di-tert-butyl dicarbonate (5.98 g, 27.41 mmol) at room temperature. The reaction mixture was stirred for several hours before it was partitioned between ethyl acetate and water. The organic layer was dried over anhydrous sodium sulfate, and then concentrated in vacuo. The residue was purified by a silica gel column chromatography eluting 5-20% ethyl acetate in hexane to afford 1,4-di(t-butoxycarbonyl)-2-(4-cyanophenyl)piperazine (2.46 g, 50.9%) as white crystals.

A solution of 1,4-di(t-butoxycarbonyl)-2-(4-cyanophenyl)piperazine (558 mg, 1.44 mmol), hydroxylamine hydrochloride (300 mg, 4.23 mmol), and sodium carbonate (763 mg, 7.20 mmol) in ethanol (3 mL) and water (3 mL) was heated at 80 °C for 2.5 h before it was partitioned between dichloromethane and water. The aqueous layer was extracted with dichloromethane. The combined organic layer was dried over sodium sulfate, and concentrated in vacuo to afford white foam (680 mg), which was dissolved in toluene (5 mL) and treated with triethyl orthoformate (2.4 mL, 14.4 mmol) and p-toluenesulfonic acid (27 mg, 0.14 mmol). The solution was heated at 90 °C for 1 h before it was partitioned between dichloromethane and saturated sodium bicarbonate aqueous solution. The organic layer was dried over anhydrous sodium sulfate, and concentrated in vacuo. The resulting white crystals were washed with ethyl acetate, and dried in vacuo to afford 1,4-di(t-butoxycarbonyl)-2-(4-(1,2,4-oxadiazol-3-yl)phenyl)piperazine (464 mg, 75% in 2 steps).

To a solution of 1,4-di(t-butoxycarbonyl)-2-(4-(1,2,4-oxadiazol-3-yl) phenyl)piperazine (464 mg, 1.08 mmol) in dichloromethane (2 mL) was added 4 N hydrogen chloride in ethyl acetate (3 mL) at room temperature. After stirring for



1.5 h, the precipitate was collected and dried in vacuo to afford 2-(4-(1,2,4-oxadiazol-3-yl)phenyl)piperazine dihydrochloride (321 mg, 98%) as a white powder.

To a suspension of 2-(4-(1,2,4-oxadiazol-3-yl)phenyl)piperazine dihydrochloride (102 mg, 0.34 mmol) in tetrahydrofuran (6 mL) was added triethylamine (0.23 mL, 1.65 mmol) and then 2-chloro-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one (73 mg, 0.33 mmol) at room temperature. After stirring for 24 h, the resulting mixture was concentrated in vacuo. The residue was dissolved in dichloromethane and saturated sodium bicarbonate aqueous solution, and the solution was passed through CHEM ELUT CE1010 (manufactured by VARIAN). The filtrate was concentrated in vacuo to yield crystals, which were washed with diisopropyl ether and ethanol to afford (S)-2-(3-(4-(1,2,4-oxadiazol-3-yl)phenyl) piperazin-1-yl)-3-methyl-6-(4-pyridyl)pyrimidin-4-one (102 mg 74%) as a white powder.

Example 18: Synthesis of 2-[4-(2-Methoxyphenylamino)-piperidin-1-yl]-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (No. XB276)

To a solution of anisidine (3.1g, 25.2 mmol) and
4-oxo-piperidine-1-carboxylic acid tert-butyl ester (5.0 g, 25.1 mmol) in methanol
(100 mL) was added sodium triacetoxyborohydride (13.4 g, 63.2 mmol) at room
temperature. After stirring for 6 h, the resulting suspension was partitioned
between ethyl acetate and 1N sodium hydroxide. The aqueous layer was extracted
with ethyl acetate. The combined organic layer was washed with brine, dried over
magnesium sulfate, and concentrated in vacuo. The residue was purified by silica
gel chromatography eluting 10-20 % ethyl acetate in hexane to furnish
4-(2-methoxyphenylamino)-piperidine-1-carboxylic acid tert-butyl ester (2.7g,
8.8mmol, 35%) as a pale yellow oil.

To a solution of 4-(2-methoxyphenylamino)-piperidine-1-carboxylic acid tert-butyl ester (2.7g, 8.8mmol) in methanol (30 mL) was added 4N hydrochloric

acid in ethyl acetate (20 mL) at room temperature. After stirring for 1h, the resulting suspension was concentrated in vacuo. The residue was partitioned between chloroform and 1N sodium hydroxide. The aqueous layer was extracted with chloroform. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel chromatography eluting 10-20% methanol in chloroform to furnish 4-(2-methozyphenylamino)-piperidine (1.8 g, 8.7 mmol, 99%) as white crystals.

To a solution of 4-(2-methoxyphenylamino)-piperidine (0.8 g, 3.87 mmol) and triethylamine (1.3 g, 12.8 mmol) in tetrahydrofuran (20 mL) was added 2-chloro-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (0.8 g, 3.61 mmol) portionwise. After stirring for 12 h, the resulting suspension was partitioned between chloroform and 1N sodium hydroxide. The aqueous layer was extracted with chloroform. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel chromatography eluting 5-10% methanol in chloroform to furnish 2-(4-(2-methoxyphenylamino)-piperidin-1-yl)-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (1.2 g, 3.07 mmol, 85%) as white crystals.

Example 19: Synthesis of 3-Methyl-2-(3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidin-1-yl)-6-(pyridin-4-yl)-3H-pyrimidin-4-one (No. XB278)

A solution of (4-bromo-phenyl)-acetic acid ethyl ester (2.31 g, 9.50 mmol) in dimethylsulfoxide (6 mL) was added to the suspension of sodium hydride (407 mg, 60% in oil, 10.17 mmol) and stirred 3 min. A solution of (3-bromo-propyl)-carbamic acid tert-butyl ester (2.03 g, 8.52 mmol) in dimethylsulfoxide (6 mL) was added to the solution and stirred at 50 °C for 30 min. The resulting solution was partitioned between ethyl acetate and saturated aqueous ammonium chloride. The aqueous layer was extracted with ethyl acetate. The combined organic layer was washed with water and brine, dried by passing through Celite column, and concentrated in





vacuo. The residue was purified by silica gel chromatography eluting ethyl acetate / hexane (4/1 to 3/1, v/v) to afford 3-(4-Bromo-phenyl)-6-tert-butoxycarbonylamino-hexanoic acid ethyl ester (2.43 g, 74%).

To a solution of 3-(4-Bromo-phenyl)-6-tert-butozycarbonylamino-hexanoic acid ethyl ester (2.43 g, 6.32 mmol) in ethyl acetate (3 mL) was added 4 N hydrogen chloride in ethyl acetate (6 mL) at room temperature. Removal of the solvent in vacuo after stirring for 30 min afforded 6-Amino-3-(4-bromo-phenyl)-hexanoic acid ethyl ester hydrochloride that was used in the next step without further purification.

A solution of 6-amino-3-(4-bromo-phenyl)-hexanoic acid ethyl ester hydrochloride, potassium carbonate (1039 mg, 7.52 mmol) in ethanol (50 ml) was refluxed for 20 hr. Solvent was removed in vacuo after addition of dilute hydrochloric acid and water was added to the residue. Filtration, wash with water and dryness afforded 3-(4-Bromo-phenyl)-piperidin-2-one (1387 mg, 86%, 2 steps).

To an ice-cooled solution of 3-(4-bromo-phenyl)-piperidin-2-one (37.97 g, 149 mmol) in tetrahydrofuran (250 ml) was added borane-tetrahydrofuran complex (335 ml, 1.0 M solution in THF, 335 mmol). The solution was stirred overnight at room temperature, and then refluxed 1.5 hr after addition of 10% aqueous hydrochloric acid. Solvents was removed in vacuo, and the residue was partitioned between dichloromethane and 1N sodium hydroxide. The aqueous layer was extracted with dichlorometane. The combined organic layer was washed with water and brine, dried over sodium sulfate, and concentrated in vacuo. The residue was dissolved in water (100 mL) and concentrated hydrochloric acid (100 mL) and refluxed for 3 hr. Sodium hydroxide was added to the solution and the resulting solution was extracted with dichlorometane. The organic layer was washed with water and brine, dried over sodium sulfate Concentration in vacuo afforded 3-(4-bromo-phenyl)-piperidine (32 18 g, 90%).

To a suspension of 3-(4-bromophenyl)-piperidine (25.2 g, 105 mmol), and

triethylamine (13 g, 128 mmol) in tetrahydrofuran (250 mL) was added di-tert-butyl-dicarbonate (25.2 g, 105 mmol) at room temperature. After stirring for 1 h, the resulting suspension was partitioned between ethyl acetate and 1N sodium hydroxide. The aqueous layer was extracted with ethyl acetate. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was washed by hexane to furnish 3-(4-bromophenyl)- piperidine-1-carboxylic acid tert-butyl ester (35.7 g, 105 mmol, 100%) as white crystals.

To a suspension of 3-(4-bromophenyl)-piperidine-1-carboxylic acid tert-butyl ester (3.0 g, 8.8 mmol), palladium acetate (80 mg, 0.36 mmol), 2-(di-t-butyl phosphino)biphenyl (210 mg, 0.70 mmol), and sodium t-butoxide (1.2 g, 125 mmol) in toluene (30 mL) was added N-methylpiperazine (1.3 g, 13.0 mmol) at room temperature. After heating at 90 °C for 5 h, the resulting suspension was passed through a Celite column. The filtrate was concentrated under reduced pressure, and the residue was purified by silica gel chromatography eluting 5-25% of ethyl acetate in hexane to afford 3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidine-1-carboxylic acid tert-butyl ester (2.0 g, 5.56 mmol, 63%) as white crystals.

To a solution of 3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidine1-carboxylic acid tert-butyl ester (2.0 g, 5.56 mmol) in methanol (20 mL) was added
4N hydrochloric acid in ethyl acetate (20 mL) at room temperature. After stirring
for 1h, the resulting suspension was concentrated in vacuo. The residue was washed
with ethyl acetate to furnish 3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidine
trihydrochloride (1.84 g, 4.99 mmol, 90%) as white crystals.

To a solution of 3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidine trihydrochloride salt (0.4 g, 1.08 mmol) and triethylamine (0.6 g, 5.93 mmol) in tetrahydrofuran (10 mL) was added 2-chloro-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (0.22 g, 0.99 mmol) portionwise. After stirring for 12 h, the

resulting suspension was partitioned between chloroform and 1N sodium hydroxide. The aqueous layer was extracted with chloroform. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel chromatography eluting 5-10% methanol in chloroform to furnish 3-methyl-2-(3-(4-(4-methylpiperazin-1-yl)-phenyl)-piperidin-1-yl)-6-(piridin-4-yl)-3H-pyrimidin-4-one (0.31 g, 0.70 mmol; 71%) as white crystals.

Example 20: Synthesis of 2-(3-(4-cyclohexylaminophenyl)-piperidin-1-yl)-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (No. XB301)

To a suspension of 3-(4-bromophenyl)-piperidine-1-carboxylic acid tert-butyl ester (8.0 g, 23.5 mmol), palladium acetate (210 mg, 0.94 mmol), 2-(di-t-butyl phosphino)biphenyl (560 mg, 1.88 mmol), and sodium t-butoxide (3.2 g, 33.3 mmol) in toluene (80 mL) was added cyclohexylamine (2.8 g, 28.2 mmol) at room temperature. After heating at 90 °C for 5 h, the resulting suspension was passed through a Celite column. The filtrate was concentrated under reduced pressure, and the residue was purified by silica gel chromatography eluting 5-25% of ethyl acetate in hexane to afford 3-(4-cyclohexylaminophenyl)-piperidine-1-carboxylic acid tert-butyl ester (6.74 g, 18.8 mmol, 80%) as white crystals.

To a solution of 3-(4-cyclohexylaminophenyl)-piperidine-1-carboxylic acid tert-butyl ester (6.74 g, 18.8 mmol) in methanol (50 mL) was added 4N hydrochloric acid in ethyl acetate (40 mL) at room temperature. After stirring for 1 h, the resulting suspension was concentrated in vacuo. The residue was washed with ethyl acetate to furnish 3-(4-cyclohexylaminophenyl)-piperidine dihydrochloride (5.84 g, 17.6 mmol, 94%) as white crystals.

To a solution of 3-(4-cyclohexylaminophenyl)-piperidine dihydrochloride salt (1.0 g, 3.02 mmol) and triethylamine (1.5 g, 14.8 mmol) in tetrahydrofuran (20

mL) was added 2-chloro-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (0.64 g, 2.89 mmol) portionwise. After stirring for 12 h, the resulting suspension was partitioned between chloroform and 1N sodium hydroxide. The aqueous layer was extracted with chloroform. The combined organic layer was washed with brine, dried over magnesium sulfate, and concentrated in vacuo. The residue was purified by silica gel chromatography eluting 5-10% methanol in chloroform to furnish 2-(3-(4-cyclohexylaminophenyl)-piperidin-1-yl)-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (1.23 g, 2.77 mmol, 96%) as white crystals.

Example 21: Synthesis of 2-(4-(4-Bromo-phenyl)-piperidin-1-yl)-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one (No. XB267)

Mixture of 4-bromobenzaldehyde (22.40 g, 121.1 mmol), dimethyl malonate(19.37 g, 146.6 mmol), cat. acetic acid and cat. piperidine in toluene (100 ml) were refluxed for 6 h with azeotropically removal of water. Resulting solution was partitioned between ethyl acetate and water. The aqueous layer was extracted with ethyl acetate. The combined organic layer was washed with water, saturated aqueous sodium bicarbonate and brine, dried over sodium sulfate. Concentration of the organic solvent in vacuo afforded 2-(4-bromo-benzylidene)-malonic acid diethyl ester as an oil that was used in the next step without further purification.

To an ice-cooled solution of dimethyl malonate (19.35 g, 146.5 mmol) and sodium methoxide (30. 12g in 28% methanol solution, 156.1 mmol) in methanol (300 ml) was added 2-(4-bromo-benzylidene)-malonic acid diethyl ester in methanol (50 ml). After stirring for 3 h, the solvent was removed in vacuo and the residue was partitioned between ethyl acetate and dilute hydrochloric acid. The aqueous layer was extracted with ethyl acetate. The combined organic layer was washed with brine, dried over sodium sulfate. Concentration of the organic solvent in vacuo afforded 3-(4-bromo-phenyl)-2,4-bis-ethoxycarbonyl-pentanedioic acid diethyl ester as an oil that was used in the next step without further purification.

A solution of 3-(4-bromo-phenyl)-2,4-bis-ethoxycarbonyl-pentanedioic acid diethyl ester in concentrated hydrochloric acid (100 ml) and acetic acid (100 ml) was refluxed for 8 h. Removal of the solvent in vacuo and recrystallization of the residue from acetonitrile yielded 3-(4-bromo-phenyl)-pentanedioic acid (22.84 g in 1st crop, 65%, 3.84 g in 2nd crop, 11.05% from 4-bromobenzaldehyde).

A solution 3-(4-bromo-phenyl)-pentanedioic acid (26.68 g, 92.9 mmol) in acetic anhydride (100 ml) was refluxed for 1.5 hr. Removal of the solvent in vacuo, and remaining solvent were azeotropically removed using toluene.

Teterahydrofuran (200 ml) and aqueous ammonia (28%, 50 ml) was added to the residue and stirred overnight. After removal of the solvent in vacuo, acetic anhydride (100 ml) was added and refluxed for 4 hr. After removal of the solvent in vacuo and succeeding azeotropic distillation with toluene, residue was partitioned between ethyl ether and water. Filtration of the suspension and dryness afforded the 4-(4-bromo-phenyl)-piperidine-2,6-dione (12.53 g, 50%) as a solid.

To an ice-cooled solution of lithium tetrahydroborate (4.13 g, 189.6 mmol) in tetrahydrofuran (200 ml) was added chlorotrimethylsilane (41.52 g, 382.2 mmol). After stirring 5 min, a solution of 4-(4-bromo-phenyl)-piperidine-2,6-dione (12.53 g, 46.7 mmol) was added and stirred overnight. The resulting solution was concentrated in vacuo after addition of 10% aqueous hydrochloric acid. The residue was dissolved in aqueous sodium hydroxide solution and methanol, and a solution of di-tert-butyl dicarbonate (11.45 g, 52.5 mmol) in methanol (10 ml) was added and stirred for 6 h. After removal of the solvent in vacuo, concentrated hydrochloric acid wad added and stirred overnight. After extraction of the solution by diethyl ether, sodium hydroxide was added to the aqueous layer to turn basic, and extracted with dichloromethane. The organic layer was washed with brine, dried over sodium sulfate. The residue of the diethyl ether and dichloromethane after removal of the solvents under reduced pressure was mixed and dissolved in tetrahydrofuran (200 ml). A solution of di-tert-butyl dicarbonate (7.45 g, 34.1 mmol) in tetrahydrofuran

WO 2004/085408

PCT/JP2004/004320

(10 ml) and triethylamine were added and stirred overnight. The resulting solution was concentrated in vacuo. Purification of the residue by silica gel chromatography eluting hexane / ethyl acetate (5/1, v/v) furnished

4-(4-bromo-phenyl)-piperidine-1-carboxylic acid tert-butyl ester (14.4g, 91%) as a solid.

To a solution of furnished 4-(4-bromophenyl)-piperidine-1-carboxylic acid tert-butyl ester (1114 mg, 3.27 mmol) in ethyl acetate (1 mL) was added 4 N hydrogen chloride in ethyl acetate (2 mL) at room temperature. After stirring for 5 h, solvent was removed in vacuo, and the resulting solid was washed with ethyl acetate and dried in vacuo to afford (4-(4-bromophenyl)-piperidine hydrochloride (884 mg, 98%) as a white solid.

A solution of (4-(4-bromophenyl)-piperidine hydrochloride (279 mg, 1.01 mmol) and triethylamine (554 mg, 5.47 mmol), 2-chloro-3-methyl-6- (pyridin-4-yl)-3H-pyrimidin-4-one (206 mg, 0.929 mmol) in tetrahydrofuran (20 mL) was stirred for 3 hr. The resulting solution was diluted with tetrahydrofuran and filtrated. After removal of the solvents under reduced pressure and the purification of the resulting residue by CHEM ELUT CE1010 (manufactured by VARIAN) eluting dichloromethane / ethanol (15/1, v/v) and wash with ethyl acetate afforded 2-(4-(4-Bromophenyl)-piperidin-1-yl)-3-methyl-6-pyridin-4-yl-3H-pyrimidin-4-one (368 mg, 93%) as a solid.

Example 22: Synthesis of 3-Methyl-6-pyridin-4-yl-2-[4-(4-pyrrolidin-1-yl-phenyl)-piperidin-1-yl]-3H-pyrimidin-4-one (No. XB269)

A suspension of 4-(4-Bromophenyl)-piperidine-1-carboxylic acid tert-butyl ester (1.97 g, 5.79 mmol), palladium acetate (54 mg, 0.24 mmol), 2-(di-t-butylphosphino)biphenyl (154 mg, 0.52 mmol), and sodium t-butoxide (846 mg, 8.80 mmol), pyrrolidine (587 mg, 8.25 mmol) in toluene (80 mL) was heated at 90 °C for 3 h under nitrogen atmosphere. The resulting suspension was passed through a

Celite column and partitioned between ethyl acetate and water. The aqueous layer was extracted with ethyl acetate. The combined organic layer was washed with brine, dried over sodium sulfate, and concentrated in vacuo. Purification of the residue by HPLC afforded 4-(4-pyrrolidin-1-yl-phenyl)-piperidine-1-carboxylic acid tert-butyl ester as a solid that was used in the next step without further purification.

To a solution of furnished 4-(4-Pyrrolidin-1-yl-phenyl)-piperidine-1-carboxylic acid tert-butyl ester in ethyl acetate (5 mL) was added 4 N hydrogen chloride in ethyl acetate (10 mL) at room temperature. After stirring for 3 h, solvent was removed in vacuo, and the resulting solid was purified by HPLC. Sodium hydroxide was added to the resulting fractions and the aqueous layer was extracted by dichloromethane. Organic layer was washed with brine, and passed through Cerite. Removal of the solvent under reduced pressure afforded 4-(4-pyrrolidin-1-yl-phenyl)-piperidine (1.01 g, 76%).

A solution of 4-(4-pyrrolidin-1-yl-phenyl)-piperidine (215 mg, 0.933 mmol) and triethylamine (391 mg, 3.86 mmol), 2-chloro-3-methyl-6-(pyridin-4-yl)-3H-pyrimidin-4-one (187 mg, 0.844 mmol) in tetrahydrofuran (10 mL) was refluxed for 5 hr. The resulting solution was diluted with tetrahydrofuran and filtrated. After removal of the solvents under reduced pressure and the purification of the resulting residue by CHEM ELUT CE1010 (manufactured by VARIAN) eluting dichloromethane / ethanol (15/1, v/v) and wash with ethyl acetate afforded 3-methyl-6-pyridin-4-yl-2-(4-(4-pyrrolidin-1-yl-phenyl)-piperidin-1-yl)-3H-pyrimidin-4-one (284 mg, 81%) as a solid.

Example 23: Synthesis of 2-(4-(6-Fluorobenzo[b]thiophen-3-yl)piperidin-1-yl)-1-methyl-1H-[4,4']bipyrimidinyl-6-one (No. YB253)

The key intermediate 4-(6-fluorobenzo[b]thiophen-3-yl)piperidine hydrochloride of 2-[4-(6-fluorobenzo[b]thiophen-3-yl)piperidin-1-yl]-1-methyl- 1H-

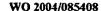
[4,4]bipyrimidinyl-6-one was synthesized from 1-acetylpipridine-4-carboxylic acid which was prepared according to the method reported by Watanabe (*J. Heterocyclic Chem.*, 30, 445 (1993)).

To a solution of 1-benzoylpiperidine-4-carbozylic acid (66 g, 285 mmol) in dichloromethane (160 mL) was added thionyl chloride (26 mL, 388 mmol). After stirring at 60°C for 1 h, the mixture was added portionwise to a stirred suspension of 2,4-difluorobenzene (45 g, 397 mmol) and anhydrous aluminum chloride (88 g, 666 mmol) in dichloromethane (245 mL), and the reaction mixture was refluxed for 5 h. The reaction mixture was poured into a mixture of ice and concentrated hydrochloric acid and extracted with chloroform. The organic layer was dried over sodium sulfate and the solvent was evaporated under reduced pressure.

Recrystallization from hexane gave 1-benzoyl-4-(2,4-difluorobenzoyl)piperidine (46 g, 50%) as colorless crystals.

A solution of 1-benzoyl-4-(2,4-difluorobenzoyl)piperidine (40 g, 120 mmol), methyl thioglycolate (12 mL, 130 mmol) in dimethylformamide (500 mL) was stirred at room temperatute for 12h. The solvent was evaporated off in vacuo and the residue treated with water and ethyl acetate. The organic layer was dried over sodium sulfate and the solvent was evaporated under reduced pressure. The obtained residue was purified by silica gel column chromatography eluting hexane/ethyl acetate to give 3-(1-benzoylpiperidin-4-yl)-6-fluorobenzo[b]thiophene-2 -carboxylic acid (11.8 g, 26%) as an oil.

3-(1-Benzoylpiperidin-4-yl)-6-fluorobenzo[b]thiophene-2-carboxylic acid (10 g, 26 mmol) was suspended in quinoline (100 mL) and cupper powder (0.5g) was added. After stirring at 200°C for 1 h, the mixture was cooled to room temperature and partitioned between ethyl acetate and water. The organic layer was dried over magnesium sulfate and evaporated. The obtained residue was purified by silica gel column chromatography eluting hexane/ ethyl acetate to give (4-(6-fluorobenzo[b]thiophen-3-yl)piperidin-1-yl)phenylmethanone (5.0 g, 48%) as yellow



crystals.

A solution of (4-(6-fluorobenzo[b]thiophen-3-yl)piperidin-1-yl) phenylmethanone (6.5 g, 19 mmol) in acetic acid (100 mL) and concentrated hydrochloric acid (100 mL) was stirred at 90°C for 10 h. To a solution of reaction mixture was added ethyl acetate. The precipitated crystals were collected by filtration and washed with ethyl acetate to give 4-(6-fluorobenzo[b]thiophen-3-yl)piperidine hydrochloride (4.8 g, 89%) as yellow crystals.

To a solution of 4-(6-fluorobenzo[b]thiophen-3-yl)piperidine hydrochloride (200 mg, 0.74 mmol) and 2-chloro-1-methyl-1H-[4,4']bipyrimidinyl-6-one (160 mg, 0.70 mmol) in tetrahydrofuran (10 mL) was added triethylamine (212 mg, 2.1 mmol). The mixture was stirred at 90°C for 6 h. The solvent was evaporated off in vacuo and the residue was treated with water and chloroform. The organic layer was dried over sodium sulfate and the solvent was evaporated under reduced pressure. Recrystallization from ethyl acetate gave 2-[4-(6-fluorobenzo[b]thiophen-3-yl)piperidin-1-yl]-1-methyl-1H-[4,4']bipyrimidinyl-6-one (220 mg, 96%) as colorless crystals.

Example 24: Synthesis of 2-(4-(Biphenyl-2-yl)piperazin-1-yl)-1-methyl-1*H*-[4,4']bipyrimidinyl-6-one (No. YA1552)

To a solution of 1-biphenyl-2-yl-piperazine dihydrochloride (311 mg, 1.0 mmol) and 2-chloro-1-methyl-1H-[4,4']bipyrimidinyl-6-one (202 mg, 0.91 mmol) in tetrahydrofuran (20 mL) was added triethylamine (404 mg, 4.0 mmol). The mixture was stirred at 90°C for 4 h. The solvent was evaporated off in vacuo and the residue treated with water and chloroform. The organic layer was dried over sodium sulfate and the solvent was evaporated under reduced pressure. Recrystallization from ethyl acetate gave 2-[4-(biphenyl-2-yl)piperazin-1-yl]-

1-methyl-1H-[4,4']bipyrimidinyl-6-one (250 mg, 65%) as colorless crystals.



The compounds in the following table were prepared in the same manner as the methods described above. The compound numbers in the following table correspond to those shown in the above-described table of preferred compounds.

Table 5

NO	NMR	Exact-MS
XA19	2.51-2.89(4H, m), 3.31-3.34(4H, m), 3.39(3H,s), 3.56(2H, s), 6.80(1H, s), 7.25-7.31(1H, m), 7.31-7.36(4H, m), 7.98(2H, dd, J=1.5, 4.8 Hz), 8.68(2H, dd, J=1.5, 4.5 Hz)(DMSO-d6)	362
XA25	3.32-3.34(4H, m), 3.46(3H, s), 3.48-3.51(4H, m), 6.80-6.85(1H, m), 6.84(1H, s), 7.01(2H, d, J=8.0 Hz), 7.23-7.28(2H, m), 8.00(2H, dd, J=1.3, 4.6 Hz), 8.70(2H, dd, J=1.5, 4.5 Hz)(DMSO-d6)	348
XA156	3.47(3H,s), 3.51-3.60(4H, m), 3.62-3.71(4H, m), 6.85(1H, s), 7.41-7.49(1H, m), 7.56-7.61(1H, m), 8.02(2H, dd, J=1.5, 4.5 Hz), 8.09(1H, d, J=8.1 Hz), 8.16(1H, d, J=8.1 Hz), 8.70(2H, dd, J=1.5, 4.8 Hz)(DMSO-d6)	405
XA289	1.11-1.28(3H, m), 2.98-3.16(1H, m), 3.28-3.41(1H, m), 3.39(3H, s), 3.54-3.80(3H, m), 3.88-3.99(1H, m), 4.08-4.26(4H, m), 4.32-4.45(1H, m), 7.13(1H, s), 7.37-7.53(5H, m), 8.45(2H, d, J=5.8 Hz), 8.96(2H, d, J=6.0 Hz) (DMSO-d6)	434
XA361	3.44(3H,s), 3.54-3.95(6H,m), 4.64(1H,brs), 7.11(1H,s), 7.42-7.51(3H,m), 7.74(2H,d,J=6.6Hz), 8.46(2H,d,J=5.7Hz), 8.94(2H,d,J=5.7Hz), 9.98(1H,brs), 10.46(1H, brs) (DMSO-d6).	348
XA364	(DMSO-d6): 3.41-3.76(4H, m), 3.48(3H, s), 3.89-4.01(2H, m), 4.96(1H, m), 7.16(1H, s), 7.33-7.58(3H, m), 8.11(1H, dd, J=7.2, 7.2Hz), 8.52(2H, d, J=6.6Hz), 8.97(2H, d, J=6.6Hz), 10.04(1H, m), 10.66(1H, m).	366
XA365	3.43(s, 3H), 3.51-3.96(m, 6H), 4.70(m, 1H), 7.00(s, 1H), 7.25(m, 1H), 7.54(m, 2H), 7.60(m, 1H), 8.20(d, J=5.7Hz, 2H), 8.80 (d, J=5.7Hz, 2H)(CDCl3)	366
XA366	2.27-2.85(1H, m), 2.94-3.08(3H, m), 3.43(3H,s), 3.59-3.67(2H, m), 3.94-3.97(1H, m), 6.81(1H, s), 7.19(2H, t, J=8.9 Hz), 7.50-7.55(2H, m), 7.96(2H, dd, J=1.6, 4.5 Hz), 8.68(2H, dd, J=1.5, 4.6 Hz)(DMSO-d6)	366



	To oc o rotott	
1	3.35-3.50(2H, m), 3.46(3H, s), 3.58-3.75(2H,	1
V	m), 3.86-3.97(2H, m), 4.68(1H, t, J=9.3 Hz),	
XA366	7.15(1H, s), 7.35(2H, t, J=9.0 Hz),	366
(HCI)	7.82-7.87(2H, m), 8.48(2H, d, J=6.6 Hz),	300
	8.96(2H, d, J=6.3 Hz), 9.55-10.08(1H, m),	
	10.54-10.70(1H, m)(DMSO-d6)	
	(CDCl3):2.81(1H,dd,J=10.4,12.5Hz),	
	3.18-3.40(3H,m), 3.50-3.80(5H,m),	l
XA369	4.50(1H,dd,J=2.5,10.1Hz), 6.67(1H,s),	
V4308	7.20-7.45(3H,m), 7.74(1H,dd,J=1.9,7.6Hz),	382
1	7.81(2H,dd,J=1.4,4.6Hz),	Ì
1	8.70(2H,dd,J=1.4,4.6Hz).	i
<u> </u>	(CDCl3):3.01(1H,dd,J=10.4,12.5Hz),	
1	3.10-3.30(3H,m), 3.50-3.80(5H,m),	ł
	4.04(1H,dd,J=2.7,10.8Hz), 6.67(1H,s),	
XA370	7.20-7.45(4H,m), 7.50(1H,s),	382
1 .		
ł	7.80(2H,dd,J=1.5,4.8Hz),	
<b></b>	8.71(2H,dd,J=1.5,5.1Hz).	
1	3.44(3H,s), 3.44-3.71(7H,m), 3.90(2H,m),	·
	4.66(1H,brs), 7.11(1H,s),	
XA371	7.55(2H,d,J=8.4Hz), 7.78(2H,d,J=8.4Hz),	382
	8.50(2H,d,J=5.7Hz), 8.95(2H,d,J=5.7Hz),	
	10.13(1H,brs), 10.60(1H,brs)(DMSO-d6)	
ļ	(DMSO-d6):3.45(3H,s), 3.50-4.20(6H,m),	
XA376	4.66(1H,br s), 7.12(1H,s), 7.72(4H,s),	426
1	8.44(2H,d,J=6.6Hz), 8.94(2H,d,J=6.6Hz),	420
	10.00(1H,br s), 10.05(1H,br s).	
	3.37-3.93(6H, m), 3.48(3H, s), 3.87(3H, s),	
Į.	4.89-4.95(1H, m), 7.04-7.12(2H, m),	
	7.17(1H, d, J=8.5 Hz), 7.45-7.51(1H, m),	
XA391	7.75-7.81(1H, m), 8.29-8.38(2H, m),	378
1	8.83-8.91(2H, m), 9.66-9.77(1H, m),	,
	0.03-0.31(21), III), 9.00-9.77(1H, III),	i
· · · · ·	9.91-10.10(1H, m)(DMSO)	
	(DMSO-d6) :3.30-3.58(5H,m),	
}	3.58-3.80(2H,m), 3.81(3H,s),	
XA392	3.85-4.00(2H,m), 4.58-4.75(1H,m),	
AA392	7.03(1H,dd,J=1.8,8.1Hz), 7.11(1H, s),	.378
	7.26(1H,d,J=7.8Hz), 7.35-7.50(2H,m),	
	8.41(2H,d,J=5.7Hz), 8.92(2H,d,J=6.0Hz),	
	9.80-10.00(1H,brd), 10.30-10.60(1H,brd).	
	3.40-3.43(5H,m), 3.51-3.63(2H,m),	
	3.78(3H,s), 3.93(2H,m),4.58(1H,br),	
XA393	7.02-7.06(3H,m), 7.64(2H,d,J=8.7Hz),	378
	8.34(2H,d,J=6.3Hz), 8.88(2H,d,J=8.7Hz).	,
	9.76(1H,br), 10.16(1H,br)(DMSO-d6)	
	1.30(3H, t, J=6.9 Hz), 3.38-3.54(1H, m),	
	3.49(3H, s), 3.65-3.79(1H, m), 3.84-3.98(2H,	
	m), 4.02-4.18(2H, m), 4.84(1H, t, J=10.5 Hz),	
XA396	7.04-7.16(2H, m), 7.15(1H, s), 7.39-7.45(1H,	202
	m), 7.89(1H, d, J=6.6 Hz), 8.49(2H, d, J=6.3	392
	Hz), 8.95(2H, d, J=6.6 Hz), 9.92(1H, d, J=9.3	
	Hz), 10.51-10.64(1H, m)(DMSO-d6)	
	, 10.01-10.04(111, 111)(DMSO-00)	



(DMSO-36);3,442H,m), 3,94(2H,J,=11,4Hz), 4,02-4,40(5H,m), 4,78(1H,J,=10,4Hz), 7.06(1H,s), 7.98(2H,d,J=8,3Hz), 8.01(2H,d,J=8,3Hz), 9.02(1H,d,J=5,1Hz), 9.31(1H,d,J=1.2Hz), 10.03(1H,d,J=8,7Hz), 10.57(1H,s), (CDCi3);2.00(4H,m), 3.03(3H,dd,J=10,8,12.0Hz), 3.21(3H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 3.90(1H,dd,J=2.7,10.8Hz), 6.57(2H,d,J=8,7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=8,7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=8,7Hz), 7.80(2H,m), 3.87(4H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8,7Hz), 7.36(2H,d,J=8,7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), (CDCi3);3.02(1H,d,J=10,712.4Hz), 3.18(7H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8,7Hz), 7.36(2H,d,J=8,7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=8,7Hz), 7.80(2H,d,J=8,7Hz), 7.80(2H,d,J=8,7Hz), 7.80(2H,d,J=8,7Hz), 8.70(2H,d,J=8,1Hz),		(DMCO 40) 0 04/5-	
XA406 7.98(2H,d,J=8.3Hz), 8.01(2H,d,J=8.3Hz), 8.23(1H,dd,J=1.2.5.1Hz), 9.02(1H,d,J=5.1Hz), 9.31(1H,d,J=1.2Hz), 10.03(1H,d,J=5.1Hz), 10.57(1H,s).  (CDC(3):2.00(4H,m), 3.57(3H,s), 3.62(2H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 3.90(1H,dd,J=2.7,10.8Hz), 6.57(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=6.8Hz), 7.29(2H,d,J=6.8Hz), 8.70(2H,d,J=6.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=6.5Hz), 8.70(2H,d,J=6.3Hz), 9.31(3H,m), 3.48(3H,m), 3.69-3.73(1H,m), 3.78(3H,m), 3.82(3H,m), 4.89(1H,t,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0Hz), 7.08(1H,d,J=9.0Hz), 9.31(1H,d,J=9.0Hz), 9.31(1H,d,J=0.5Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=0.3Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=0.3Hz), 8.96(2H,d,J=6.3Hz), 10.07(1H,m), 8.30(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 9.70(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 9.70(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 9.97(2H,m), 3.88(3H,s), 4.81-4.91(1H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-4.00(6H,m), 3.60-4.00(6H,m), 3.46(3H,s), 3.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(2H,d,J=6.8Hz), 8.94(2H,d,J=6.8Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(2H,d,J=6.8Hz), 7.14(1H,m), 8.95(2H,d,J=6.8Hz), 7.49-5.80(1H,m), 8.44-8.50(2H,d,J=6.8Hz), 7.49-5.80(1H,m), 8.44-8.50(2H,d,J=6.8Hz), 7.49-5.80(1H,m), 8.44-8.50(2H,d,J=6.8Hz), 7.49-5.80(1H,m), 8.44-8.50(2H,d,J=6.8Hz), 7.49-5.80(1H,m), 8.44-8.50(2H,d,J=6.8Hz), 7.49-5.80(1		(DMSO-d6):3.64(2H,m),	
7.56(2F1,d,J=6.3Hz), 8.01(2H,d,J=8.3Hz), 9.02(1H,d,J=5.1Hz), 9.02(1H,d,J=5.1Hz), 9.31(1H,d,J=1.2Hz), 10.03(1H,d,J=8.7Hz), 10.57(1H,s).  (CDCI3):2.00(4H,m), 3.03(1H,dd,J=10.8,12.0Hz), 3.21(3H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 417  (5.7(2H,d,J=8.7Hz), 6.66(1H,s), 7.29(2H,d,J=8.7Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 9.18(7H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.20(1H,L,S), 13.81(1H,L,S), 13.81(1H,LS),	XA406	4.78(1H t.l=10.4Hz), 7.06(4H, m),	
9.02(HH,d,J=5.HHz), 9.31(HH,d,J=1.2Hz), 10.03(HH,d,J=8.7Hz), 10.57(1H,s), (CDCI3):2.00(4H,m), 3.03(HH,dd,J=10.8,12.0Hz), 3.21(3H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 3.90(1H,dd,J=2.7,10.8Hz), 6.57(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), (CDCI3):3.02(HH,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), (CDCI3):2.36(3H,s), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.25(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=6.3Hz), 8.96(2H,d,J=4.5Hz), 1.70(2H,d,J=4.5Hz), 8.70(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 8.91(H,d,J=0.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0 Hz), 7.15(1H,s), 7.66(1H,d,J=6.3 Hz), 9.93(1H,d,J=0.9Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 8.97(2H,m), 3.40-3.50(4H,m), 3.47(3H,s), 3.83-3.94(2H,m), 3.88(3H,s), 4.81-4.91(1H,m), 6.92-6.99(1H,m), 7.07-7.01(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 8.50-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.64-0.06(1,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.85-7.00(1H,m), 7.95(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 8.40(2H,d,J=6.8Hz), 8.52-8.64(1H,m), 8.46(2H,d,J=6.8Hz), 8.52-8.64(1H,m), 8		6   7.98(2H.d.J=8.3Hz)   8.04(2H.d.J=8.3Hz)	
9.02(1H,d,J=5.1Hz), 9.31(1H,d,J=1.2Hz), 10.03(1H,d,J=8.7Hz), 10.57(1H,s).  (CDCIS):2.00(4H,m), 3.03(1H,dd,J=10.8,12.0Hz), 3.21(3H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 4.17  (5.7(2H,d,J=8.7Hz), 6.66(1H,s), 7.29(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.18(7H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 7.36(2H,d,J=4.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.20(1H,H,d),=1.6Hz), 3.22(7H,m), 3.02(1H,L,J=11.6Hz), 3.22(7H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 9.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 9.94(1H,d,J=0.9Hz), 7.15(1H,s), 7.80(2H,d,J=4.5Hz), 8.96(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.80(4H,m), 8.30(2H,m), 8.30-8.40(2H,m), 8.80(3H,m), 3.80(3H,s), 3.89(3H,s), 4.81-4.91(1H,m), 8.30-8.40(2H,m), 8.80(3H,m), 3.80(3H,s), 8.96(3H,d), 3.80(3H,s), 3.89(3H,s), 4.81-4.91(1H,m), 8.30-8.40(2H,m), 8.80-8.92(2H,m), 9.70-9.79(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 8.30-8.40(2H,m), 8.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80-6.00(6H,m), 3.60-3.80(1H,m), 3.80(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 8.96(2H,d,J=6.8Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 7.95(1H,d,J=6.6Hz), 8.94(2H,d,J=6.8Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m		8.23(1H.dd.J=1.2.5 1Hz)	373
(CDCI3):2.00(4H,m), 3.03(1H,dd,J=10.81,2.0Hz), 3.21(3H,m), 3.29(4H,m), 3.57(3H,s), 3.62(2H,m), 417 6.57(2H,d,J=8.7Hz), 6.66(1H,s), 7.29(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), (CDCI3):3.02(1H,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=4.8Hz), (CDCI3):2.36(3H,s), 2.59(4H,m), 3.02(1H,d,J=4.5Hz), 7.36(2H,d,J=4.5Hz), (CDCI3):2.36(3H,s), 2.59(4H,m), 3.02(1H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), (CDCI3):2.36(3H,s), 2.59(4H,m), 3.02(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.73(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(	1	9.02(1H,d,J=5.1Hz), 9.31(1H,d, 1=4.2H=)	
(CDCI3):2.00(H4,m), 3.03(H1,dd,J=10,8,12.0Hz), 6.57(2H,d,J=8,7Hz), 6.66(1H,s), 7.29(2H,d,J=8,7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), (CDCI3):3.02(1H,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.99(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), (CDCI3):2.36(3H,s), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=8,7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 7.80(2H,d,J=8,7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=8,7Hz), 7.34(2H,d,J=8.7Hz), 7.80(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8,7Hz), 7.34(2H,d,J=8.7Hz), 7.80(1H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0Hz), 7.15(1H,s), 7.66(1H,d,J=3.0Hz), 8.51(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=0.0Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.20(1H,m), 8.30(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 10.07(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 3.40-3.50(4H,m), 3.40(3H,s), 3.83-3.94(2H,m), 8.85-8.92(2H,m), 9.70-9.79(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 8.466-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.	ļ	10.03(1H,d,J=8.7Hz), 10.57(1H s)	- 1
XA433  3.90(H,dd,J=2.7,10.8Hz), 6.57(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), (CDCl3):3.02(H,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=4.8Hz), (CDCl3):2.36(3H,S), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.94(2H,d,J=4.5Hz), 8.94(2H,d,J=4.5Hz), 8.94(3H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0 Hz), 7.15(1H,s), 7.66(1H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=9.0 Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.90(1H,m), 8.92(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 9.97(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 7.12(1H,s), 7.97-91(1H,m), 7.09(1H,d,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,d,J=6.8Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.8Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.8Hz), 7.14(1H,s), 7.49-7.58(1H,m), 8.44-8.50(2H,m), 8.52-8.64(1H,m), 8.96(2H,d,J=6.8Hz), 7.44-7.58(1H,m), 8.96(2H,d,J=6.8Hz), 8.96(2H,d		(CDCl3):2.00(4H m)	<del></del> -
XA433  3.90(H,dd,J=2.7,10.8Hz), 6.57(2H,d,J=8.7Hz), 7.80(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), (CDCl3):3.02(H,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=4.8Hz), (CDCl3):2.36(3H,S), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.94(2H,d,J=4.5Hz), 8.94(2H,d,J=4.5Hz), 8.94(3H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0 Hz), 7.15(1H,s), 7.66(1H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=9.0 Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.90(1H,m), 8.92(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 9.97(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 7.12(1H,s), 7.97-91(1H,m), 7.09(1H,d,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,d,J=6.8Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.8Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.8Hz), 7.14(1H,s), 7.49-7.58(1H,m), 8.44-8.50(2H,m), 8.52-8.64(1H,m), 8.96(2H,d,J=6.8Hz), 7.44-7.58(1H,m), 8.96(2H,d,J=6.8Hz), 8.96(2H,d	1	3.03(1H,dd,J=10.8,12.0Hz), 3.21(3H m)	1
3.50(1H,ad,J=2.7,10.8Hz), 6.57(2H,d,J=8.7Hz), 6.66(1H,s), 7.29(2H,d,J=8.7Hz), 6.66(1H,s), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 8.70(2H,d,J=4.8Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.29(4H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.36(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 9.70(1H,m), 7.80(1H,d,J=9.0 Hz), 7.15(1H,s), 7.66(1H,d,J=3.0 Hz), 9.93(1H,d,J=9.0 Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 9.93(1H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 10.07(1H,m), 8.20(1H,m), 8.30(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 10.07(1H,m), 7.97.91(1H,m), 7.97.91(1H,m), 8.30(2H,d,J=6.3Hz), 8.87(2H,d,J=6.9e(91H,m), 7.07-7.10(1H,m), 7.92(1H,m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-6.7.00(1H,brd), 10.35-10.60(1H,brd), 10.	VA 400	1 4.49(411,111), 3.37(3H.S) 3 62/2H m/	
7.29(2H,d,J=8.7Hz), 6.66(1H,s), 8.70(2H,d,J=4.8Hz).  (CDCl3):3:02(1H,dd,J=10.7,12.4Hz), 3.18(7H,m), 3.55(3H,s), 3.62(2H,m), 3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz).  (CDCl3):2:36(3H,s), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 3.41-3.54(3H,m), 3.48(3H,s), 3.69-3.73(1H,m), 3.78(3H,s), 3.82(3H,s), 3.69-3.73(1H,m), 3.78(3H,s), 3.82(3H,s), 3.86-3.93(2H,m), 4.89(1H,t,J=10.5Hz), 6.97-7.01(1H,m), 7.08(1H,d,J=9.0Hz), 7.15(1H,s), 7.66(1H,d,J=6.3Hz), 9.93(1H,d,J=9.0Hz), 10.60-10.73(1H,m)(DMSO-d6)  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.96(2H,d,J=6.3Hz), 10.07(1H,m), 3.40-3.50(4H,m), 8.30(2H,d,J=6.3Hz), 8.87(2H,d), 3.88(3H,s), 4.81-4.91(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 0.02-10.23(1H,m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.9,8,7Hz), 8.46(2H,d,J=6.6Hz), 7.14(1H,s), 7.49-7.58(1H,m), 8.96(2H,d,J=6.8Hz), 8.52-8.64(1H,m), 8.96(2H,d,J=6.8Hz)	AA433	,   2.80(1H'00'1=5'\'.10'8H <sup>2</sup> /	417
XA439  XA4439  XA443  XA442  XA442  XA442  XA442  XA444  XA444  XA444  XA444  XA445  XA445  XA446  XA468  X		6.57(2H,d,J=8.7Hz), 6.66(1H,s),	
XA439  XA4439  XA443  XA442  XA442  XA442  XA442  XA444  XA444  XA444  XA444  XA445  XA445  XA446  XA468  X	1	7.29(2H,d,J=8./Hz), 7.80(2H,d,J=4.8Hz),	
XA439  3.87(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 8.96(2H,d,J=6.3Hz), 9.93(1H,d,J=9.0 Hz), 7.05(1H,m), 7.06(1H,m), 7.08(1H,d,J=9.0 Hz), 7.15(1H,s), 7.66(1H,d,J=6.3 Hz), 9.93(1H,d,J=9.0 Hz), 10.60-10.73(1H,m)(DMSO-d6)  XA464  XA464  (DMSO-d6): 3.45(3H,s), 3.38-3.81(6H,m), 3.88(6H,s), 5.06(1H,m), 6.82(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,d,J=6.3Hz), 8.87(2H,m), 3.88(3H,s), 4.81-4.91(1H,m), 6.92-6.99(1H,m), 7.07-7.10(1H,m), 7.12(1H,s), 7.79-7.91(1H,m), 8.30-8.40(2H,m), 8.85-8.92(2H,m), 9.70-9.79(1H,m), 3.60-3.80(1H,m), 3.60-3.80(1H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 3.60-3.80(1H,m), 8.85-8.92(2H,m), 9.70-9.79(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.46(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H,m), 3.44(3H,s), 3.94(3H,s), 4.94-5.02(1H,m), 6.96-7.01(1H,m), 7.05(1H,d,J=6.6Hz), 8.94(2H,d,J=6.8Hz), 7.49-7.58(1H,m), 8.44-8.50(2H,m), 8.52-8.64(1H,m), 8.96(2H,d,J=6.6Hz), 7.49-7.58(1H,m), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 7.49-7.58(1H,m), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.6Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,d,J=6.Hz), 8.96(2H,	<del> </del>	O.70(2H,U,J=4.8HZ)_	
XA439 S.01(4H,m), 3.96(1H,dd,J=2.5,11.1Hz), 6.66(1H,S), 6.93(2H,d,J=8.7Hz), 7.36(2H,d,J=8.7Hz), 7.79(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz).  (CDCi3):2.36(3H,s), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 3.41-3.54(3H, m), 3.48(3H, s), 3.69-3.73(1H, m), 3.78(3H, s), 3.82(3H, s), 3.69-3.73(1H, m), 4.89(1H, t, J=10.5 Hz), 6.97-7.01(1H, m), 7.08(1H, d, J=9.0 Hz), 7.15(1H, s), 7.66(1H, d, J=6.3 Hz), 8.96(2H, d, J=6.3Hz), 8.97(2H, d, J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m), 3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.98.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	1	3 18/7H m) 3 55/3H a) 3 20/2Hz),	
XA463  XA464  XA464  XA464  XA464  XA468  XA468  XA468  XA468  XA469/  XA470  XA470  AB	VA 400	3.87(4H m) 3.96(1H dd 1=2.544.41)	-
XA442  XA442  XA442  XA442  XA444  XA4444  XA444  XA444  XA444  XA444  XA444  XA444  XA444  XA444  XA4444  XA44444	AA439	6.66(1H.S), 6.93(2H.d. l=0.7H=)	434
XA442  XA442  XA442  XA442  XA444  XA	1	7.36(2H,d,J=8.7Hz) 7.79(2H,d,J=4.5H=)	
(CDCl3):2.36(3H,s), 2.59(4H,m), 3.02(1H,t,J=11.6Hz), 3.22(7H,m), 3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz). 3.41-3.54(3H, m), 3.48(3H, s), 3.69-3.73(1H, m), 3.78(3H, s), 3.82(3H, s), 3.69-3.73(1H, m), 3.78(3H, s), 3.82(3H, s), 3.69-3.73(1H, m), 4.89(1H, t, J=10.5 Hz), 6.97-7.01(1H, m), 7.08(1H, d, J=9.0 Hz), 7.15(1H, s), 7.66(1H, d, J=3.0 Hz), 8.51(2H, d, J=6.3 Hz), 8.96(2H, d, J=6.3 Hz), 9.93(1H, d, J=9.0 Hz), 10.60-10.73(1H, m)(DMSO-d6)  (DMSO-d6): 3.45(3H, s), 3.38-3.81(6H, m), 3.88(6H, s), 5.06(1H, m), 6.82(2H, d, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m), 3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m), 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9).8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd). 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.46-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		8.70(2H,d,J=4.5Hz),	
XA442  3.55(3H,s), 3.63(2H,m), 3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 3.41-3.54(3H, m), 3.48(3H, s), 3.69-3.73(1H, m), 3.78(3H, s), 3.82(3H, s), 3.69-3.73(1H, m), 4.89(1H, t, J=10.5 Hz), 6.97-7.01(1H, m), 7.08(1H, d, J=9.0 Hz), 7.15(1H, s), 7.66(1H, d, J=3.0 Hz), 8.51(2H, d, J=6.3 Hz), 8.96(2H, d, J=6.3 Hz), 9.93(1H, d, J=9.0 Hz), 10.60-10.73(1H, m)(DMSO-d6)  XA464  XA464  XA464  XA464  XA464  XA464  XA468  XA469/ XA470  XA469/ XA470  XA469/ XA470  XA469/ XA470  XA469/ XA470  XA469/ XA470	1	(CDCl3):2.36(3H.s) 2.59(4H m)	<del> </del>
XA442  3.94(1H,d,J=10.5Hz), 6.66(1H,s), 6.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz), 7.80(2H,d,J=4.5Hz), 8.70(2H,d,J=4.5Hz), 3.41-3.54(3H, m), 3.48(3H, s), 3.69-3.73(1H, m), 3.78(3H, s), 3.82(3H, s), 3.69-3.73(1H, m), 4.89(1H, t, J=10.5 Hz), 6.97-7.01(1H, m), 7.08(1H, d, J=9.0 Hz), 7.15(1H, s), 7.66(1H, d, J=3.0 Hz), 8.51(2H, d, J=6.3 Hz), 8.96(2H, d, J=6.3 Hz), 9.93(1H, d, J=9.0 Hz), 10.60-10.73(1H, m)(DMSO-d6)   (DMSO-d6): 3.45(3H, s), 3.38-3.81(6H, m), 3.88(6H, s), 5.06(1H, m), 6.82(2H, d, J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m), 3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.98.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.95-6.64(1H, m), 8.96(2H, d, J=6.6 Hz)	1	3.02(1H,t,J=11.6Hz), 3.22(7H m)	
XA464  XA464  XA464  XA468  XA468  XA468  XA468  XA468  XA469/  XA469/  XA469/  XA469/  XA469/  XA469/  XA472  XA474  XA4	XA442	3.55(3H,S), 3.63(2H,m)	
XA463  XA464  XA464  XA464  XA464  XA464  XA464  XA464  XA464  XA468  XA469/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA470		3.94(1H,d,J=10.5Hz), 6.66(1H,s),	446
XA463  XA464  XA464  XA464  XA464  XA464  XA464  XA464  XA464  XA468  XA469/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA460/  XA470		0.93(2H,d,J=8.7Hz), 7.34(2H,d,J=8.7Hz),	
XA463  XA463  XA463  XA463  XA464  XA466  XA468  XA469/  XA469/  XA469/  XA470  XA469/  XA470  XA469/  XA470  XA469/  XA470  XA469/  XA470  XA	<b> </b>	<u>」,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,</u>	
XA463  XA463  XA463  XA464  XA466  XA468  XA469  XA469  XA469  XA470  XA470  XA469  XA470  XA	l	m) 3 78(3H s) 3 82(3H s) 3 69-3.73(1H,	
XA463  A, J=3.0 Hz), 8.51(2H, d, J=6.3 Hz), 8.96(2H, d, J=6.3 Hz), 9.93(1H, d, J=9.0 Hz), 10.60-10.73(1H, m)(DMSO-d6)  (DMSO-d6): 3.45(3H, s), 3.38-3.81(6H, m), 3.88(6H, s), 5.06(1H, m), 6.82(2H, d, J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m).  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 3.60-3.80(1H, m) (DMSO)  (DMSO-d6): 3.38-3.60(6H, m), 3.60-3.80(1H, m), 3.80-4.00(5H, m), 4.80-4.97(1H, m), 6.85-7.00(1H, m), 7.95(1H, dd, J=6.9Hz), 8.94(2H, d, J=6.3Hz), 9.80-10.00(1H, brd), 10.35-10.60(1H, brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.96(2H, d, J=6.6 Hz), 9.96(2H, d, J=6.6 Hz), 9.96(2H, d, J=6.6 Hz), 9.96(2H	]	m), 4.89(1H, 1, 1=10, 5 Hz), 6.07, 7.04(41)	1 1
XA464  XA466  XA466  XA466  XA466  XA466  XA468  XA469  XA469  XA469  XA469  XA469  XA469  XA469  XA469  XA469  XA470  XA469  XA470  XA	XA463	7.08(1H, d, J=9.0 Hz), 7.15(1H, e), 7.65(4H,	
XA464  XA464  (DMSO-d6): 3.45(3H, s), 3.38-3.81(6H, m), 3.88(6H, s), 5.06(1H, m), 6.82(2H, d, J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H, m), 3.60-3.80(1H, m), 3.80-4.00(5H, m), 4.80-4.97(1H, m), 6.85-7.00(1H, m), 7.95(1H, dd, J=2.4, 11.4Hz), 7.13(1H, s), 7.95(1H, dd, J=6.6Hz), 8.94(2H, d, J=6.3Hz), 9.80-10.00(1H, brd), 10.35-10.60(1H, brd), 10.35-10.60(1H, brd), 10.35-10.60(1H, brd), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz), 3.96(2H, d, J=6.6 Hz), 3.96(2H, d, J=6.6 Hz), 8.96(2H, d, J=6.6 Hz), 3.96(2H, d, J=6.6 Hz), 3.96(		1 4, 0 3, 0 112), 0.51(2H, 0, J=6 3 Hz) 8 06/20	408
XA464  XA464  (DMSO-d6): 3.45(3H, s), 3.38-3.81(6H, m), 3.88(6H, s), 5.06(1H, m), 6.82(2H, d, J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m).  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H, m), 3.60-3.80(1H, m); 8.80-4.97(1H, m), 6.85-7.00(1H, m), 7.95(1H, dd, J=2.4, 11.4Hz), 7.13(1H, s), 7.95(1H, dd, J=6.98.7Hz), 8.46(2H, d, J=6.6Hz), 8.94(2H, d, J=6.3Hz), 9.80-10.00(1H, brd), 10.35-10.60(1H, brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	1	1 4, 3-6.3 HZ), 9.93(1H, d. J=9 0 H <sub>2</sub> )	ł
XA464  J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m).  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m); 3.60-3.80(1H,m); 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		10.60-10.73(1H, m)(DMSO-d6)	}
XA464  J=8.7Hz), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m).  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m); 3.60-3.80(1H,m); 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		(DMSO-d6): 3 45/3H s) 3 29 2 94/6H	
XA464  3-3.7+2), 7.04(1H, s), 7.44(1H, t, J=8.4Hz), 8.20(1H, m), 8.30(2H, d, J=6.3Hz), 8.87(2H, d, J=6.3Hz), 10.07(1H, m).  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		3.88(6H, s), 5.06(1H, m), 6.82(2H, d)	1 1
XA469/ XA469/ XA470  XA472  XA472  XA472  XA472  XA472  XA472  XA472  XA472  XA472  XA468  XA468  XA468  XA468  XA468  XA468  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	XA464	1 3-0.7 (TZ), 7.04(1H, S), 7.44(1H + 1=8 AU-)	1 400
XA468  XA469  XA469  XA469  XA469  XA469  XA470  XA469  XA470  XA		1 0.20(17, m), 8.30(2H, d, J=6 3Hz) 8 87/24	408
XA468  3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H, m), 3.88(3H, s), 4.81-4.91(1H, m), 6.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		[ 4, 3-0.3H2), 10.07(1H, m).	1
XA468		3.40-3.50(4H, m), 3.47(3H, s), 3.83-3.94(2H	<del></del>
XA468  C.92-6.99(1H, m), 7.07-7.10(1H, m), 7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd), 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		1 "" 7, 0.00(311, S), 4.01-4.91(1H m)	]
7.12(1H, s), 7.79-7.91(1H, m), 8.30-8.40(2H, m), 8.85-8.92(2H, m), 9.70-9.79(1H, m), 10.02-10.23(1H, m)(DMSO)  (DMSO-d6): 3.38-3.60(6H,m), 3.60-3.80(1H,m) 3.80-4.00(5H,m), 4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	XA468	0.92-6.99(1H, m), 7.07-7.10/1H m)	
XA469/ XA470  XA472  XA473  XA474  XA474  XA474  XA474  XA475  XA476  XA477  XA477  XA477  XA477  XA477  XA477  XA477  XA477  XA478  X	,	1.12(1H, s), 7.79-7.91(1H, m) 8 30-8 40(2H)	396
XA469/ XA469/ XA470  XA470  (DMSO-d6):3.38-3.60(6H,m), 3.60-3.80(1H,m):3.80-4.00(5H,m), 4.80-4.97(1H,m):6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz);7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz);8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd);10.35-10.60(1H,brd); 3.36-4.00(6H, m);3.46(3H, s);3.94(3H, s), 4.94-5.02(1H, m);6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz);7.14(1H, s), 7.49-7.58(1H, m);8.44-8.50(2H, m), 8.52-8.64(1H, m);8.96(2H, d, J=6.6 Hz)		1117, 0.05-0.92(2H, m), 9.70-9.79/1H m)	·
XA469/ XA470  XA470  X		(DMSO-d6) (2.29 2.00(01)	
XA469/ XA470  4.80-4.97(1H,m), 6.85-7.00(1H,m), 7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd). 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		3.60-3.88(1H m) 3.80.4.00(5H, m)	
XA470  7.09(1H,dd,J=2.4,11.4Hz), 7.13(1H,s), 7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd).  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	YAACOL	4.80-4.97(1H m) 6.85-7.00(4H m)	
7.95(1H,dd,J=6.9,8.7Hz), 8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd). 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		7.09(1H,dd,J=2.4.11.4Hz) 7.12(4H.a)	
8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.3Hz), 9.80-10.00(1H,brd), 10.35-10.60(1H,brd). 3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	AA410	7.95(1H,dd,J=6.9.8.7Hz)	396
XA472  XA472  3.36-4.00(6H, m), 3.46(3H, s), 3.94(3H, s), 4.94-5.02(1H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)		8.46(2H,d,J=6.6Hz), 8.94(2H,d,J=6.2H=)	
XA472  XA473  XA473  XA474  XA474  XA475  XA475  XA475  XA476  XA476  XA477  XA		9.80° 10.00(1H,Drd), 10.35-10 60(1H hed)	i
XA472 (7H, m), 6.96-7.01(1H, m), 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	]	3.50-4.00(6H, m), 3.46(3H s) 3 94/3H s	
XA472 7.05(1H, d, J=8.6 Hz), 7.14(1H, s), 7.49-7.58(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, J=6.6 Hz)	l	4.34-3.02(1H, m), 6.96-7.01/1H m)	
8.52-8.64(1H, m), 8.44-8.50(2H, m), 8.52-8.64(1H, m), 8.96(2H, d, l=6.6 Hz)	XA472	1.00(1H, d, J=8.6 Hz), 7.14(1H e)	200
10.49-10.60(1H, m)(DMSO)		1.45-1.58(1H, m), 8.44-8.50(2H m)	396
1.0.43-10.00(1H, M)(DMSO)	l	10.49-10.60/14>/01/20/20	. [
		10.40-10.00(1H, M)(DMSO)	





XA480	2.78(1H, dd, J=10.0, 12.1 Hz), 3.18-3.27(3H, m), 3.59(3H, s), 3.64-3.74(2H, m), 3.86(3H, s), 4.37(1H, dd, J=2.4, 10.1 Hz), 6.67(1H, s), 6.89(1H, d, J=2.1 Hz), 6.99(1H, dd, J=1.7, 8.0 Hz), 7.50(1H, d, J=8.2 Hz), 7.82(2H, dd, J=1.5, 4.8 Hz), 8.71(2H, dd, J=1.8, 4.5 Hz)(CDCI3)	412
XA490 (2HCI)	3.35-3.94(6H, m), 3.49(3H, s), 4.71-4.80(1H, m), 7.02-7.11(1H, m), 7.18-7.28(2H, m), 7.98-8.10(1H, m), 8.31-8.48(2H, m), 8.87-8.97(2H, m), 9.79-9.92(1H, m), 10.18-10.39(1H, m) (DMSO)	380
XA501	(CDCl3):2.77(1H,dd,J=10.2,12.0Hz), 3.15-3.35(3H,m), 3.50-3.80(5H,m), 3.84(3H,s), 4.39(1H,d,J=7.8Hz), 6.67(1H,s), 6.78(1H,d,J=8.8Hz), 7.39(1H,dd,J=2.4,8.7Hz), 7.71(1H,d,J=2.3Hz), 7.82(2H,d,J=6.0Hz), 8.71(2H,d,J=6.0Hz).	456
XA510	(CDCl3): 1.98-2.05(4H, m), 2.85(1H, dd, J=12, 10.5Hz), 3.17-3.24(7H, m), 3.58(3H, s), 3.65-3.72(2H, m), 3.85(3H, s), 4.28(1H, dd, 10.5, 2.7Hz), 6.10(1H, d, J=2.1Hz), 6.18(1H, dd, J=8.7, 2.1Hz), 6.65(1H, s), 7.33(1H, d, J=8.4Hz), 7.83(2H, dd, J=4.5, 1.8Hz), 8.70(2H, dd, J=4.5, 1.5Hz).	447
XA511	(CDCl3):1.90-2.05(4H,m), 2.93(1H,t,J=12.0Hz), 3.15-3.40(7H,m), 3.59(3H,s), 3.65-3.85(5H,m), 4.11(1H,dd,J=2.1,10.2Hz), 6.49(1H,dd,J=3.0,9.0Hz), 6.66(1H,s), 7.83(2H,dd,J=1.8,4.5Hz), 8.70(2H,dd,J=1.5,4.5Hz).	447
XA516	(DMSO-d6):3.20-3.70(4H,m), 3.70(1H,m), 3.98(3H,s), 3.99(3H,s), 4.00(1H,m), 4.96(1H,d,J=10.2Hz), 7.01(1H,s), 7.03(2H,m), 8.26(2H,d,J=6.1Hz), 8.53(1H,s), 8.84(2H,d,J=6.1Hz), 10.25(1H,d,J=10.7Hz)	414
XA525	(DMSO-d6):3.30-3.50(2H,m), 3.48(3H,s), 3.55-3.78(2H,m), 3.78(3H,s), 3.96(2H,d,J=13.5Hz), 4.69(1H,t,J=10.4Hz), 7.06(1H,t,J=7.4Hz), 7.12(1H,s), 7.14(1H,d,J=7.4Hz), 7.31(1H,d,J=7.4Hz), 7.39(1H,t,J=7.4Hz), 7.59(2H,d,J=8.3Hz), 7.77(2H,d,J=8.3Hz), 8.43(2H,d,J=6.5Hz), 8.93(2H,d,J=6.5Hz), 9.89(1H,d,J=8.7Hz), 10.34(1H,s).	454
XA527	(DMSO-d6):3.40-4.10(9H,m), 3.81(3H,s), 4.69(1H,m), 7.05(1H,s), 7.05(2H,d,J=9.0Hz), 7.67(2H,d,J=9.0Hz), 7.75(4H,s), 8.27(2H,d,J=5.7Hz), 8.85(2H,d,J=5.7Hz), 9.75(1H,s), 10.04(1H,s).	454



	Transport	
XA536	(DMSO-d6):3.40-3.60(2H,m), 3.47(3H,s), 3.68(2H,m), 3.95(2H,m), 4.71(1H,t,J=9.9Hz), 7.16(1H,s), 7.33(2H,t,J=8.85Hz), 7.78(6H,m), 8.50(2H,d,J=6.3Hz), 8.97(2H,d,J=6.3Hz), 10.02(1H,s), 10.50(1H,s).	443
XA543	3.52(s, 3H), 3.57-4.10(m, 6H), 5.57(m, 1H), 7.02(s, 1H), 7.53-7.70(m, 2H), 8.06(d, J=7.2Hz, 2H), 8.21-8.34(m, 3H), 8.82(d, J=6.3Hz, 2H), 9.88-9.92(m, 1H), 10.58-10.61(m, 1H)(DMSO d6)	398
XA544	3.41-3.59(2H, m), 3.49(3H, s), 3.68-3.76(2H, m), 3.97-4.02(2H, m), 4.78-4.89(1H, m), 7.15(1H, s), 7.58-7.63(2H, m), 7.89-8.07(4H, m), 8.30(1H, s), 8.49(2H, d, J=6.3 Hz), 8.95(2H, d, J=6.3 Hz), 10.17(1H, d, J=8.4 Hz), 10.57-10.70(1H, m)(DMSO-d6)	398
XA619	(CDCl3): 2.98(1H, dd, J=12.6, 10.8Hz), 3.17-3.28(5H, m), 3.58(3H, s), 3.62(1H, m), 3.79(1H, m), 4.26(1H, dd, 10.5, 2.7Hz), 4.62(2H, m), 6.66(1H, s), 6.88(1H, t, J=7.5Hz), 7.16(1H, d, J=7.2Hz), 7.27(1H, m), 7.84(2H, d, J=6.0), 8.70(2H, dd, J=4.8, 1.2Hz).	390
XA626	3.33-3.41(4H, m), 3.42(3H, s), 3.47-3.87(4H, m), 6.84(1H, s), 7.44-7.49(5H, m), 7.99(2H, dd, J=1.5, 4.5 Hz), 8.69(2H, dd, J=1.4, 4.8 Hz)(DMSO-d6)	376
XA649	3.44(3H, s),3.37-4.04(9H, m),4.67(1H, d,J=9.6Hz),7.10(1H, s),7.45-7.55(3H, m),7.83(2H, d,J=6.0Hz),8.47(2H, d,J=6.6Hz),8.95(2H, d,J=6.6Hz),12.15(1H, brs)(DMSO-d6)	362
XA756	(CDCl3):2.50-2.61(1H,m), 2.80-2.95(1H,m), 3.05-3.20(1H,m), 3.25-3.40(1H,m), 3.50-3.60(1H,m), 3.57(3H,s), 3.65-3.75(1H,m), 3.75-3.80(1H,m), 3.85(3H,s), 6.60-6.80(3H,m), 7.47(1H,dd,J=7.2,8.4Hz), 7.82(2H,dd,J=1.5,4.5Hz), 8.71(2H,dd,J=1.5,4.5Hz).	410
XA757/ XA758	(DMSO-d6):2.54(3H,s), 3.40-3.79(3H,m), 3.46(3H,s), 3.80-4.10(6H,m), 4.83-5.10(1H,m), 6.90-7.05(1H,m), 7.08(1H,s), 7.13(1H,dd,J=2.7,11.4Hz), 8.00-8.25(1H,brd), 8.37(2H,d,J=6.3Hz), 8.91(2H,d,J=6.6Hz), 11.80-12.20(1H,brd).	410
XA831	2.55(s, 3H), 3.51(s, 3H), 3.67-3.82(m, 4H), 4.04-4.08(m, 2H), 5.64(m, 1H), 7.05(s, 1H), 7.59-7.72(m, 3H), 8.06-8.11(m, 2H), 8.35(d, J=6.6Hz, 2H), 8.41(d, J=7.8Hz, 1H), 8.49 (d, J=6.9Hz, 1H), 8.84(d, J=6.6Hz, 2H)(DMSO d6)	412





	(DMSO-d6) :3.15-3.35(1H,m),	T
XA	3.38-3.60(4H,m), 3.75-4.15(8H,m),	1
1016	4.18-4.25(1H,m), 4.90-5.20(1H,m),	486
	7.00-7.20(3H,m), 7.30-7.55(6H,m),	·
	8.50-8.70(3H,m), 9.00(2H,d,J=6.3Hz).	
	(CDCI3):1.80-2.42(3H, m), 3.08-3.39(4H,	
<b>V</b> 0	m), 3.40-3.62(1H, m), 3.65-4.23(6.8H m)	1
XA	[4.63-4.90(0.6H, m), 5.40-5.62(0.7H m)	420
1276	5.80-6.00(0.1H, m), 6.52-6.78(3H, m),	438
	6.90-7.2(1H, m), 7.68-7.90(2H,m),	1
	8.64-8.80(2H,m)	<u> </u>
	1.48(3H, s), 1.57(3H, s), 3.50(3H, s),	
	3.51-3.66(2H, m), 3.72-3.76(1H, m),	1
XA	3.90(3H, s), 3.99(1H, d, J=13.4 Hz),	1
1649	5.15-5.23(1H, m), 7.08-7.12(2H, m),	406
1049	7.18(1H, d, J=8.6 Hz), 7.46-7.49(1H, m),	406
	8.04-8.11(1H, m), 8.37-8.45(2H, m),	İ
•	8.89-8.97(2H, m), 9.49-9.60(1H, m),	
	9.95-10.11(1H, m)(DMSO)	
	3.01 (1H, dd, J = 10.8, 12.9 Hz), 3.10-3.30	
XA	(3H, m), 3.50-3.75 (5H, m), 4.04 (1H, dd, J =	
1973	2.7, 10.8 Hz), 6.67 (1H, s), 7.20-7.40 (4H,	382
	m), 7.50 (1H, s), 7.80 (2H, dd, J = 1.5, 4.8	
	Hz), 8.71 (2H, dd, J = 1.5, 5.1 Hz) (CDCl3)	1
	2.80 (1H, dd, J = 10.3, 12.2 Hz), 3.15-3.30	<del></del>
	(3H, m), 3.50-3.80 (5H, m), 4.44 (1H, dd, $l = 1$	ĺ
XA	2.6, 10.3 Hz), 6.67 (1H, s), 7.10-7.20 (1H	
1974	$  m \rangle$ , 7.25-7.40 (1H, m), 7.58 (1H, dd, $J = 1.0$	426
	$  0.1 \text{ Hz} \rangle$ , $/./3$ (1H, dd, $J = 1.6.7.8 \text{ Hz} \rangle$ , $7.81 \text{ J}$	.20
	(2H, dd, J = 1.6, 4.5 Hz), 8.70 (2H, dd, J =	
	11.6, 4.5 Hz) (CDCl3)	
	2.95-3.10 (1H, m), 3.10-3.35 (3H, m), 3.56	
XA	(3H, s), 3.60-3.70 (2H, m), 3.80-4.05 (7H, m)	
1975	0.07 (1H, s), 6,87 (1H, d, J = 8.1 Hz).	407
_	6.90-7.10 (2H, m), 7.80 (2H, dd, J = 1.8, 6.3)	
	[HZ], 8.71 (2H, dd, $J = 1.5$ , 4.8 Hz) (CDC13)	
	3.40 (3H, m), 3.45 (3H, s), 3.53-3.96 (3H, m)	
XA	14.08 (1H, t, J = 13.5Hz), 7.10 (1H, s), 7.60	
1976	(2H, d, J=8.3Hz), 7.76 (2H, d, J=8.3Hz), 8.38	382
1370	(1H, Drs), 8.91 (1H, d, J=4.8Hz), 9.88 (1H, hr	302
	10) 40 24 (411 ) 10	
	s), 10.31 (1H, br s) (DMSO-d6)	
	1_	
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd	
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H t	
XA	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18/1H	
XA 1977	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d	382
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d	382
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H,	382
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) (DMSO-d6)	382
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) (DMSO-d6)  2.98 (1H, t, J = 10.9 Hz), 3.22 (m, 3H), 3.56	382
	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) (DMSO-d6)  2.98 (1H, t, J = 10.9 Hz), 3.22 (m, 3H), 3.56 (3H, s), 3.60 (2H, m), 4.03 (1H, d, J=8.7 Hz)	382
1977 XA	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) (DMSO-d6)  2.98 (1H, t, J = 10.9 Hz), 3.22 (m, 3H), 3.56 (3H, s), 3.60 (2H, m), 4.03 (1H, d, J = 8.7 Hz), 6.68 (1H, s), 7.28 (1H, d, J = 8.2 Hz), 7.46	382
1977	3.40(3H, m), 3.46(3H, s), 3.62(1H, dd, J=12.0, 13.2Hz), 3.72(1H, m), 3.92(1H, t, J=15.5Hz), 4.68(1H, t, J=10.1Hz), 7.18(1H, s), 7.58(2H, d, J=8.6Hz), 7.83(2H, d, J=8.6Hz), 8.57(2H, d, J=6.6Hz), 9.01(2H, d, J=6.6Hz), 10.20(1H, d, J=7.8Hz), 10.76(1H, br s) (DMSO-d6)  2.98 (1H, t, J = 10.9 Hz), 3.22 (m, 3H), 3.56 (3H, s), 3.60 (2H, m), 4.03 (1H, d, J=8.7 Hz)	382





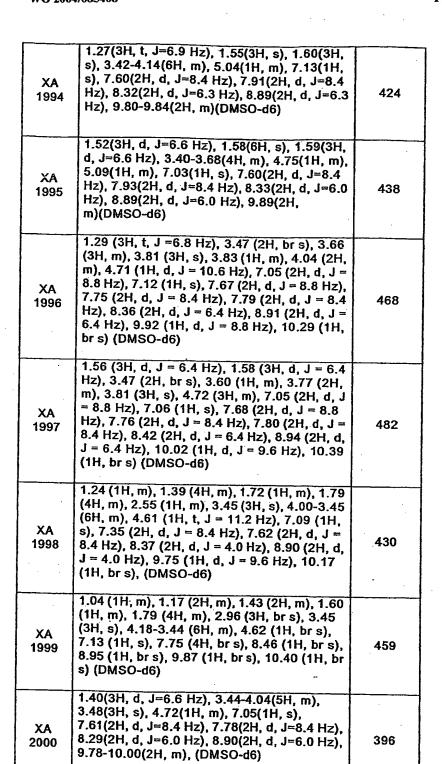
XA 1979	3.31 (1H, dd, J = 13.8, 8.9 Hz), 3.46 (3H, s), 3.85 (1H, dd, J = 13.8, 3.6 Hz), 4.10 (1H, d, J = 17.7 Hz), 4.19 (1H, d, J = 17.7 Hz), 4.91 (1H, dd, J = 8.9, 3.6 Hz), 6.11 (1H, s), 6.74 (1H, s), 7.32 (2H, d, J = 8.4 Hz), 7.42 (2H, d, J = 8.4 Hz), 7.79 (2H, dd, J = 4.8, 1.5 Hz), 8.74 (2H, dd, J = 4.8, 1.5 Hz), 6.74 (2H, dd, J = 4.8, 1.5 Hz)	396
XA 1980	1.97 (4H, m), 3.26 (4H, m), 3.38 (2H, m), 3.45 (3H, s), 3.60 (2H, m), 3.80 (1H, d, J = 13.8 Hz), 3.92 (1H, d, J = 14.1 Hz), 4.48 (1H, t, J = 10.4 Hz), 6.65 (2H, d, J = 8.7 Hz), 7.16 (1H, s), 7.54 (2H, d, J = 8.7 Hz), 8.57 (2H, d, J = 6.6 Hz), 9.00 (2H, d, J = 6.6 Hz), 9.83 (1H, d, J = 9.3 Hz), 10.32 (1H, br s) (DMSO-d6)	417
XA 1981	3.21 (4H, m), 3.40 (2H, m), 3.46 (3H, s), 3.65 (2H, m), 3.78 (4H, m), 3.91 (2H, t, J = 13.7 Hz), 4.55 (1H, t, J = 10.1 Hz), 7.14 (2H, d, J = 8.7 Hz), 7.20 (1H, s), 7.64 (2H, d, J = 8.7 Hz), 8.60 (2H, d, J = 6.6 Hz), 9.02 (2H, d, J = 6.6 Hz), 9.93 (1H, d, J = 9.0 Hz), 10.47 (1H, br s) (DMSO-d6)	433
XA 1982	2.80 (3H, d, J = 4.5 Hz), 3.15 (4H, m), 3.44 (4H, m), 3.45 (3H, s), 3.60 (2H, m), 3.82 (1H, d, J = 13.5 Hz), 3.90 (3H, m), 4.54 (1H, t, J = 10.5), 7.10 (2H, d, J = 8.7 Hz), 7.17 (1H, s), 7.64 (2H, d, J = 8.7 Hz), 8.54 (2H, d, J = 6.3 Hz), 8.99 (2H, d, J = 6.3 Hz), 9.94 (1H, d, J = 8.7 Hz), 10.47 (1H, br s), 11.26 (1H, br s) (DMSO-d6)	446
XA 1983	1.27(3H, t, J=6.6 Hz), 3.46-4.14(8H, m), 4.70(1H, m), 7.11(1H, s), 7.60(2H, d, J=8.4 Hz), 7.76(2H, d, J=8.4 Hz), 8.32(2H, d, J=6 Hz), 8.89(2H, d, J=6.0 Hz), 9.87(1H, m), 10.23(1H, m), (DMSO-d6)	396
XA 1984	1.27(6H, dd, J=6.9, 6.9 Hz), 3.37-4.36(6H, m), 4.66-4.79(2H, m), 7.03(1H, s), 7.62(2H, d, J=8.7 Hz), 7.78(2H, d, J=8.7 Hz), 8.33(2H, d, J=6 Hz), 8.90(2H, d, J=6.0 Hz), 9.93(1H, m), 10.25(1H, m), (DMSO-d6)	410
XA 1985	1.40(3H, d, J=6.3 Hz), 3.44-4.04(5H, m), 3.48(3H, s), 4.69(1H, m), 7.08(1H, s), 7.60(2H, d, J=8.4 Hz), 7.79(2H, d, J=8.4 Hz), 8.33(2H, d, J=6.3 Hz), 8.90(2H, d, J=6.3 Hz), 9.83(1H, m), 10.00(1H, m), (DMSO-d6)	396



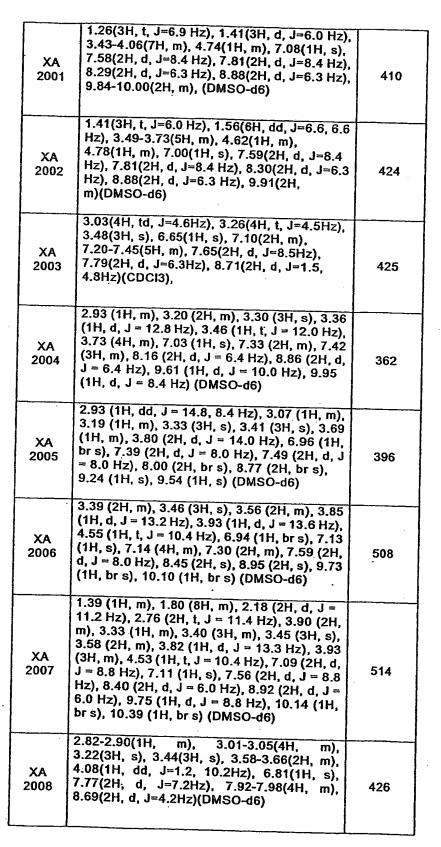


Γ	4 57(C) -> 2 50(0)	
XA 1986	1.57(6H, s), 3.50(3H, s), 3.51-3.93(4H, m), 4.98(1H, m), 7.11(1H, s), 7.60(2H, d, J=7.4 Hz), 7.94(2H, d, J=7.4 Hz), 8.41(2H, d, J=6.0 Hz), 8.93(2H, d, J=6.0 Hz), 9.88(1H, m), 10.05(1H, m), (DMSO-d6)	410
XA 1987	1.43(3H, d, J=6.6 Hz), 3.38-3.93(5H, m), 3.48(3H, s), 4.72(1H, m), 7.12(1H, s), 7.59(2H, d, J=8.4 Hz), 7.84(2H, d, J=8.4 Hz), 8.43(2H, d, J=6.6 Hz), 8.95(2H, d, J=6.6 Hz), 9.65(1H, m), 10.23(1H, m), (DMSO-d6)	396
XA 1988	2.34 (1H, m), 2.42 (1H, m), 2.80 (3H, d, J = 5.6 Hz), 2.81 (3H, d, J = 5.6 Hz), 3.28 (1H, q, J = 8.8 Hz), 3.43 (2H, m), 3.45 (3H, s), 3.57 (5H, m), 3.80 (1H, d, J = 11.4 Hz), 3.96 (2H, m), 4.50 (1H, t, J = 10.4 Hz), 6.69 (2H, d, J = 8.4 Hz), 7.14 (1H, s), 7.55 (2H, d, J = 8.4 Hz), 8.47 (2H, d, J = 5.6 Hz), 8.96 (2H, d, J = 5.6 Hz), 9.75 (1H, d, J = 8.0 Hz), 10.16 (1H, br s), 11.40 (1H, br s) (DMSO-d6)	460
XA 1989	1.65 (2H, br s), 1.91 (4H, br s), 3.46 (9H, s), 3.70 (2H, m), 3.92 (2H, t, J = 16.6 Hz), 4.66 (1H, br s), 7.16 (1H, s), 7.85 (4H, br s), 8.50 (2H, d, J = 6.4 Hz), 8.97 (2H, d, J = 6.4 Hz), 10.01 (1H, br s), 10.59 (1H, br s) (DMSO-d6)	431
XA 1990	2.32 (1H, m), 2.42 (1H, m), 2.79 (3H, d, J = 5.2 Hz), 2.81 (3H, d, J = 5.2 Hz), 3.27 (1H, m), 3.39 (2H, m), 3.45 (3H, s), 3.59 (5H, m), 3.79 (1H, d, J = 13.3 Hz), 3.95 (2H, m), 4.50 (1H, t, J = 11.6 Hz), 6.69 (2H, d, J = 8.4 Hz), 7.16 (1H, s), 7.56 (2H, d, J = 8.4 Hz), 8.50 (2H, s), 8.98 (2H, d, J = 5.6 Hz), 9.78 (1H, br s), 10.19 (1H, br s), 11.44 (1H, br s) (DMSO-d6)	460
XA 1991	3.47 (3H, s), 3.61 (3H, m), 3.81 (3H, s), 4.02 (3H, m), 4.69 (1H, t, J = 10.6 Hz), 7.05 (2H, d, J = 8.8 Hz), 7.10 (1H, s), 7.67 (2H, d, J = 8.8 Hz), 7.77 (4H, s), 8.38 (2H, br s), 8.91 (2H, d, J = 5.2 Hz), 9.90 (1H, br s), 10.28 (1H, br s) (DMSO-d6)	454
XA 1992	1.26(3H, t, J=6.9 Hz), 1.41(3H, d, J=6.3 Hz), 3.43-4.06(7H, m), 4.74(1H, m), 7.09(1H, s), 7.58(2H, d, J=8.4 Hz), 7.84(2H, d, J=8.4 Hz), 8.32(2H, d, J=6.6 Hz), 8.90(2H, d, J=6.6 Hz), 9.90(1H, m), 10.03(1H, m), (DMSO-d6)	410
XA 1993	1.41(3H, t, J=6.3 Hz), 1.55(6H, dd, J=6.6, 6.6 Hz), 3.49-3.73(5H, m), 4.64(1H, m), 4.78(1H, m), 6.99(1H, s), 7.58(2H, d, J=8.7 Hz), 7.81(2H, d, J=8.7 Hz), 8.28(2H, d, J=6.3 Hz), 8.87(2H, d, J=6.3 Hz), 9.91(2H, m)(DMSO-d6)	· 424





## WO 2004/085408







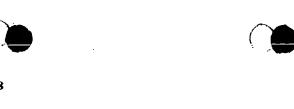
XA 2009	1.21(3H, d, J=6.6 Hz), 3.17-3.45(4H, m), 3.52(3H, s), 4.02(1H, m), 4.69(1H, m), 7.20(1H, s), 7.54(2H, d, J=8.4 Hz), 7.70(2H, d, J=8.4 Hz), 8.26(2H, d, J=6.3 Hz), 8.88(2H, d, J=6.3 Hz), 9.90(1H, m), 10.16(1H, m), (DMSO-d6)	396
XA 2010	1.21(3H, d, J=6.0 Hz), 3.17-3.45(4H, m), 3.53(3H, s), 4.02(1H, m), 4.70(1H, m), 7.24(1H, s), 7.54(2H, d, J=8.7 Hz), 7.73(2H, d, J=8.7 Hz), 8.33(2H, d, J=5.7 Hz), 8.93(2H, d, J=5.7 Hz), 10.04(1H, m), 1037(1H, m), (DMSO-d6)	396
XA 2011	3.02 (1H, t, J = 11.9 Hz), 3.17 (6H, m), 3.55 (3H, s), 3.63 (2H, m), 3.86 (4H, m), 3.96 (1H, d, J = 10.2 Hz), 6.66 (1H, s), 6.92 (2H, d, J = 8.4 Hz), 7.35 (2H, d, J = 8.4 Hz), 7.80 (2H, d, J = 5.1 Hz), 8.70 (2H, d, J = 5.1 Hz) (CDCI3)	433
XA 2012	2.31 (3.6H, s), 3.16 (4H, t, J = 4.8 Hz), 3.44 (3H, s), 3.45 (4H, m), 3.75 (4H, t, J = 4.8 Hz), 3.86 (1H, d, J = 14.0 Hz), 3.92 (1H, d, J = 12.4 Hz), 4.56 (1H, d, J = 10.4 Hz), 6.95 (1H, s), 7.06 (2H, d, J = 8.8 Hz), 7.43 (2H, d, J = 8.8 Hz), 8.06 (2H, d, J = 6.0 Hz), 8.75 (2H, d, J = 6.0 Hz), 9.03 (1H, s), 9.33 (1H,d, J = 10.0 Hz) (DMSO-d6)	433
XA 2013	1.82 (4H, m), 1.97 (2H, m), 2.12 (2H, m), 2.77 (2H, t, J = 11.6 Hz), 3.01 (2H, m), 3.27 (1H, m), 3.40 (2H, m), 3.45 (3H, s), 3.49 (2H, m), 3.57 (1H, m), 3.63 (1H, m), 3.84 (1H, d, J = 13.6 Hz), 3.92 (3H, d, J = 12.8 Hz), 4.53 (1H, t, J = 11.2 Hz), 7.12 (2H, d, J = 8.4 Hz), 7.14 (1H, s), 7.58 (2H, d, J = 8.9 Hz), 8.49 (2H, d, J = 5.2 Hz), 8.97 (2H, d, J = 5.2 Hz), 9.82 (1H, br s), 10.24 (1H, br s), 11.12 (1H, br s) (DMSO-d6)	500
XA 2014	1.75(2H, m), 2.14(2H, m), 2.72(6H, d, J=4.5 Hz), 2.74-2.80(3H, m), 3.30-3.95(8H, m), 3.45(3H, s), 4.54(1H, m), 7.10(2H, d, J=9.0 Hz), 7.15(1H, s), 7.60(2H, d, J=9.0 Hz), 8.51(2H, d, J=6.6 Hz), 8.98(2H, d, J=6.6 Hz), 9.86(1H, m), 10.32(1H, m), 10.93(1H, m), (DMSO-d6)	474
XA 2015	1.68(2H, m), 2.09(2H, m), 3.16-3.90(10H, m), 3.45(3H, s), 4.60(1H, m), 7.13(1H, s), 7.45-7.71(4H, m), 8.45(2H, d, J=6.0 Hz), 8.94(2H, d, J=6.0 Hz), 9.83(1H, m), 10.22(1H, m) (DMSO-d6)	447



WO 2004/085408



XA 2016	1.91-2.03(2H, m), 3.09(1H, m), 3.28-3.57(7H, m), 3.40(3H, s), 4.41(2H, m), 6.58(2H, d, J= 8.7 Hz), 7.13(1H, s), 7.46(2H, d, J= 8.7 Hz), 8.44(2H, d, J= 6.3 Hz), 8.94(2H, d, J= 6.3 Hz), 9.61(1H, m), 9.89(1H, m) (DMSO-d6)	433
XA 2017	2.97 (6H, s), 3.45 (3H, s), 4.20-3.30 (6H, m), 4.53 (1H, t, J = 9.8 Hz), 6.69 (2H, br s), 7.14 (1H, s), 7.57 (2H, br s), 8.48 (2H, br s), 8.96 (2H, br s), 9.72 (1H, br s), 10.09 (1H, br s) (DMSO-d6)	1
XA 2018	3.18-3.22(1H, m), 3.44-3.80(15H, m), 4.51-4.55(1H, m), 5.11(2H, s), 7.04-7.07(3H, m), 7.32-7.39(5H, m), 7.52-7.55(2H, m), 8.33-8.35(2H, m), 8.82-8.87(2H, m), 9.65-9.75(2H, br)(DMSO-d6)	566
XA 2019	1.32(6H, d, J=6.8Hz), 3.04-3.88(18H, m), 4.52-4.55(1H, m), 7.09-7.12(3H, m), 7.62(2H, d, J=7.2Hz), 8.45(2H, d, J=4.2Hz), 8.94(2H, d, J=4.2Hz), 9.83-10.34(3H, br), 11.00-11.04(1H, br)(DMSO-d6)	474
XA 2020	1.32(6H, d, J=6.8Hz), 3.04-3.88(18H, m), 4.52-4.55(1H, m), 7.09-7.12(3H, m), 7.62(2H, d, J=7.2Hz), 8.45(2H, d, J=4.2Hz), 8.94(2H, d, J=4.2Hz), 9.83-10.34(3H, br), 11.00-11.04(1H, br)(DMSO-d6)	476
XA 2021	2.09(3H, s), 3.19-4.00(20H, m), 4.43-4.54(3H, m), 7.06-7.19(3H, m), 7.62(2H, d, J=7.2Hz), 8.44(2H, d, J=4.2Hz), 8.94(2H, d, J=4.2Hz), 9.82-9.85(1H, br), 10.26-10.30(1H, br), 11.30-11.40(1H, br)(DMSO-d6)	518
XA 2022	3.17-3.21(4H, m), 3.38-4.16(14H, m), 4.51-4.54(1H, m), 7.08-7.18(3H, m), 7.60(2H, d, J=7.2Hz), 8.43(2H, d, J=4.2Hz), 8.93(2H, d, J=4.2Hz), 9.26-9.34(2H, br), 9.81-84(1H, br), 10.25-10.30(1H, br)(DMSO-d6)	432
XA 2023	1.82(3H, m), 3.29(3H, m), 3.40-3.96(9H, m), 3.48(3H, s), 4.55(1H, m), 7.10(1H, s), 7.13(2H, d, J=8.4 Hz), 7.56(2H, d, J=8.4 Hz), 8.39(2H, d, J=6.0 Hz), 8.91(2H, d, J=6.0 Hz), 9.67(1H, m), 9.97(1H, m) (DMSO-d6)	445



XA 2024	1.89-2.03(2H, m), 2.95-3.07(5H, m), 3.29-3.83(5H, m), 3.40(3H, s), 4.40(1H, m), 4.94(1H, m), 6.49(2H, d, J= 8.4 Hz), 7.13(1H, s), 7.25(2H, d, J= 8.4 Hz), 7.95(2H, d, J=6.0 Hz), 8.69(2H, d, J=6.0 Hz) (DMSO-d6)	433
XA 2025	1.16(6H, d, J= 6.3 Hz), 2.28-2.36(2H, m), 2.97-3.21(6H, m), 3.54(3H, s), 3.55-3.62(4H, m), 3.95(1H, m), 6.66(1H, s), 6.93(2H, d, J= 8.7 Hz), 7.32(2H, d, J= 8.7 Hz), 7.80(2H, d, J=6.3 Hz), 8.70(2H, d, J=6.3 Hz) (CDCI3)	460
XA 2026	1.26(6H, d, J= 6.3 Hz), 2.42(2H, dd, J= 11.1, 11.1 Hz), 3.02(1H, dd, J= 12.3, 10.8 Hz), 3.17-3.22(3H, m), 3.45-3.63(4H, m), 3.55(3H, s), 3.81(1H, m), 3.95(1H, dd, J= 13.2, 2.1 Hz), 6.66(1H, s), 6.92(2H, d, J= 8.4 Hz), 7.34(2H, d, J= 8.4 Hz), 7.80(2H, d, J=6.0 Hz), 8.70(2H, d, J=6.0 Hz) (CDCI3)	461
XA 2027	2.91-3.09(5H, m), 3.26(3H, s), 3.46(3H, s), 3.69-3.73(2H, m), 4.07-4.11(1H, m), 6.81(1H, s), 7.64(2H, d, J=7.2Hz), 7.77(2H, d, J=7.2Hz), 7.94-8.02(6H, m), 8.68(1H, d, J=4.2Hz)(DMSO-d6)	502
XA 2028	3.28-3.32(4H, m), 3.46(3H, s), 3.86-3.91(2H, m), 4.59-4.61(1H, m), 6.90(1H, s), 7.77-8.06(10H, m), 8.70(2H, d, J=4.2Hz), 9.36-9.44(1H, br)(DMSO-d6)	449
XA 2029	3.08 (1H, dd, J = 12.4, 10.4 Hz), 3.25 (3H, m), 3.58 (3H, s), 3.68 (2H, m), 4.09 (1H, dd, J = 10.4, 2.4 Hz), 6.68 (1H, s), 7.29 (2H, d, J = 8.4 Hz), 7.54 (2H, d, J = 8.4 Hz), 7.56 (2H, d, J = 8.4 Hz), 7.81 (2H, dd, J = 4.4, 1.6 Hz) (CDCI3)	·508
XA 2030	3.08 (1H, dd, J = 12.4, 10.4 Hz), 3.27 (3H, m), 3.58 (3H, s), 3.70 (2H, m), 4.11 (1H, dd, J = 10.4, 2.4 Hz), 6.68 (1H, s), 7.57 (2H, d, J = 8.0 Hz), 7.63 (2H, d, J = 8.0 Hz), 7.70 (4H, s), 7.81 (2H, dd, J = 4.8, 1.2 Hz), 8.71 (2H, dd, J = 4.8, 1.2 Hz) (CDCl3)	492
XA 2031	1.45 (3H, t, J = 12.4 Hz), 3.08 (1H, dd, J = 12.4, 10.8 Hz), 3.24 (3H, m), 3.57 (3H, s), 3.67 (2H, m), 4.07 (1H, m), 4.09 (2H, q, J = 7.0 Hz), 6.67 (1H, s), 6.97 (2H, d, J = 8.4 Hz), 7.51 (2H, d, J = 8.4 Hz), 7.51 (2H, d, J = 8.4 Hz), 7.54 (2H, d, J = 8.4 Hz), 7.57 (2H, d, J = 8.4 Hz), 7.81 (2H, dd, J = 4.8, 1.2 Hz), 8.71 (2H, dd, J = 4.8, 1.2 Hz), 8.71	468





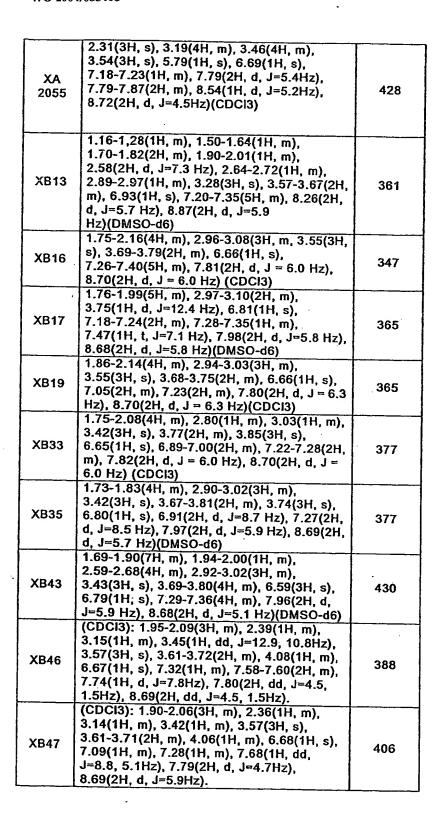
XA 2032	1.94 (4H, m), 2.02 (1H, m), 2.21 (1H, m), 2.62 (4H, m), 2.91 (1H, m), 3.03 (1H, dd, J = 12.4, 10.4 Hz), 3.20 (4H, m), 3.33 (1H, m), 3.48 (2H, m), 3.54 (3H, s), 3.62 (2H, m), 3.91 (1H, dd, J = 10.4, 2.4 Hz), 6.55 (2H, d, J = 8.4 Hz), 6.66 (1H, s), 7.29 (2H, d, J = 8.4 Hz), 7.81 (2H, dd, J = 4.4, 0.8 Hz), 8.70 (2H, dd, J = 4.4, 0.8 Hz) (CDCI3)	468
XA 2033	2.29(3H, s), 3.06(4H, t, J=4.8Hz), 3.38(4H, t, J=4.8Hz), 3.51(3H, s), 5.70(1H, s), 6.67(1H, s), 7.24-7.29(5H, m), 7.83(2H, dd, J=1.6, 4.3Hz), 8.72(2H, dd, J=1.3, 4.5Hz)(CDCI3)	427
XA 2034	3.09 (1H, dd, J = 12.0, 10.8 Hz), 3.23 (3H, m), 3.57 (3H, s), 3.66 (2H, m), 3.82 (3H, s), 3.86 (3H, s), 4.06 (1H, dd, J = 10.8, 2.4 Hz), 6.58 (2H, m), 6.67 (1H, s), 7.24 (2H, m), 7.47 (2H, d, J = 8.0 Hz), 7.53 (2H, d, J = 8.0 Hz), 7.82 (2H, dd, J = 4.8, 1.2 Hz), 8.71 (2H, dd, J = 4.8, 1.2 Hz) (CDCI3)	484
XA 2035	3.08 (3H, dd, J = 12.4, 10.8 Hz), 3.25 (3H, m), 3.57 (3H, s), 3.67 (2H, m), 3.93 (3H, s), 3.96 (3H, s), 4.08 (1H, dd, J = 10.0, 2.0 Hz), 6.68 (1H, s), 6.95 (1H, d, J = 8.4 Hz), 7.11 (1H, d, J = 2.4 Hz), 7.16 (1H, dd, J = 8.4, 2.4 Hz), 7.51 (2H, d, J = 8.0 Hz), 7.58 (2H, d, J = 8.0 Hz), 7.81 (2H, dd, J = 4.8, 1.2 Hz), 8.71 (2H, dd, J = 4.8, 1.2 Hz), (CDCl3)	484
XA 2036	3.08 (1H, dd, J = 12.4, 10.8 Hz), 3.26 (3H, m), 3.57 (3H, s), 3.67 (2H, m), 4.09 (1H, dd, J = 10.0, 2.0 Hz), 6.68 (1H, s), 7.42 (2H, d, J = 8.4 Hz), 7.53 (4H, d, J = 8.4 Hz), 7.58 (2H, d, J = 8.4 Hz), 7.80 (2H, dd, J = 4.8, 1.6 Hz), 8.71 (2H, dd, J = 4.8, 1.6 Hz) (CDCI3)	458
XA 2037	3.09 (1H, dd, J = 12.4, 10.8 Hz), 3.25 (3H, m), 3.58 (3H, s), 3.69 (2H, m), 4.11 (1H, dd, J = 10.4, 2.4 Hz), 6.68 (1H, s), 7.28 (2H, m), 7.44 (2H, d, J = 8.0 Hz), 7.51 (3H, m), 8.81 (2H, dd, J = 4.0, 1.2 Hz), 8.72 (2H, dd, J = 4.0, 1.2 Hz) (CDCI3)	492
XA 2038	3.07 (1H, dd, J = 12.3, 11.0 Hz), 3.26 (3H, m), 3.57 (3H, s), 3.67 (2H, m), 4.10 (1H, dd, J = 10.2, 2.1 Hz), 6.68 (1H, s), 7.42 (1H, dd, J = 8.1, 2.2 Hz), 7.55 (5H, m), 7.68 (1H, d, J = 2.2 Hz), 7.80 (2H, dd, J = 4.8, 1.3 Hz), 8.71 (2H, dd, J = 4.8, 1.3 Hz) (CDCl3)	492



XA 2039	3.06 (1H, dd, J = 12.0, 10.8 Hz), 3.24 (3H, m), 3.58 (3H, s), 3.67 (2H, m), 4.13 (1H, dd, J = 10.4, 2.4 Hz), 6.68 (1H, s), 7.61 (2H, d, J = 8.4 Hz), 7.80 (2H, d, J = 4.4 Hz), 8.15 (2H, d, J = 8.4 Hz), 8.71 (2H, d, J = 4.4 Hz), 8.77 (1H, s) (CDCl3)	416
XA 2040	3.04-3.26(4H, m), 3.57(3H, s), 3.66-3.71(2H, m), 4.07(1H, m), 5.12(2H, s), 6.68(1H, s), 7.06(2H, d, J= 8.7 Hz), 7.40-7.59(11H, m), 7.81(2H, d, J=6.0 Hz), 8.71(2H, d, J=6.0 Hz) (CDCI3)	530
XA 2041	0.38(2H, m), 0.67(2H, m), 1.32(1H, m), 3.09(1H, dd, J=12.6, 11.1 Hz), 3.22-3.28(3H, m), 3.58(3H, s), 3.67-3.71(2H, m), 3.86(2H, d, J= 6.9 Hz), 4.08(1H, m), 6.68(1H, s), 7.06(2H, d, J= 9.0 Hz), 7.49-7.60(6H, m), 7.82(2H, d, J=6.0 Hz), 8.72(2H, d, J=6.0 Hz) (CDCl3)	494
XA 2042	1.37(6H, d, J= 6.0 Hz), 3.08(1H, dd, J=12.3, 11.1 Hz), 3.20-3.28(3H, m), 3.57(3H, s), 3.65-3.70(2H, m), 4.06(1H, m), 4.59(1H, m), 6.67(1H, s), 7.06(2H, d, J= 9.0 Hz), 7.48-7.59(6H, m), 7.81(2H, d, J=6.0 Hz), 8.71(2H, d, J=6.0 Hz) (CDCI3)	482
XA 2043	0.99(3H, t, J= 7.5 Hz), 1.40-1.85(4H, m), 3.05-3.30(4H, m), 3.57(3H, s), 3.65-3.70(2H, m), 4.00-4.10(3H, m), 6.67(1H, s), 6.97(2H, d, J= 8.7 Hz), 7.50-7.56(6H, m), 7.81(2H, d, J=6.0 Hz), 8.71(2H, d, J=6.0 Hz) (CDCl3)	496
XA 2044	1.66(1H, br.s), 2.52(3H, s), 3.05(1H, dd, J=10.5, 12.6Hz), 3.20-3.26(3H, m), 3.57(3H, s), 3.62-3.72(2H, m), 4.07(1H, dd, J=2.1, 10.5Hz), 6.67(1H, s), 7.33(2H, d, J=8.4Hz), 7.50-7.61(6H, m), 7.81(2H, dd, J=1.6, 4.3Hz), 8.70(2H, dd, J=1.3, 4.5Hz)(CDCl3)	469
XA 2045	1.72(1H, br.s), 2.40(3H, s), 2.98-3.26(5H, m), 3.57(3H, s), 3.57-3.67(1H, m), 4.07(1H, dd, J=2.1, 10.5Hz), 6.67(1H, s), 7.24(2H, d, J=8.1Hz), 7.49-7.52(4H, m), 7.60(2H, d, J=8.1Hz), 7.81(2H, dd, J=1.6, 4.3Hz), 8.70(2H, dd, J=1.3, 4.5Hz)(CDCl3)	437
XA 2046	1.36(9H, s), 1.72(1H, br.s), 3.06(1H, dd, J=10.5, 12.4Hz), 3.20-3.28(3H, m), 3.57(3H, s), 3.57-3.67(2H, m), 4.07(1H, dd, J=2.1, 10.5Hz), 6.67(1H, s), 7.24(2H, d, J=8.1Hz), 7.43-7.56(6H, m), 7.81(2H, dd, J=1.6, 4.3Hz), 8.71(2H, dd, J=1.3, 4.5Hz)(CDCl3)	479



XA 2047	1.29(6H, d, J=6.9Hz), 1.73(1H, br.s), 2.96(1H, m), 3.06(1H, dd, J=10.5, 12.4Hz), 3.21-3.29(3H, m), 3.57(3H, s), 3.62-3.71(2H, m), 4.07(1H, dd, J=2.1, 10.5Hz), 6.67(1H, s), 7.31(2H, d, J=8.1Hz), 7.45-7.54(4H, m), 7.63(2H, d, J=8.1Hz), 7.81(2H, dd, J=1.6, 4.3Hz), 8.71(2H, dd, J=1.3, 4.5Hz), (CDCI3)	465
XA 2048	1.68(2H, br.s), 2.98(1H, dd, J=10.5, 12.6Hz), 3.20-3.27(2H, m), 3.56(3H, s), 3.64-3.74(1H, m), 4.04(1H, dd, J=3.3, 11.1Hz), 4.80(3H, s), 6.66(1H, s), 6.72(2H, d, J=8.5Hz), 7.49-7.52(4H, m), 7.63(2H, d, J=8.1Hz), 7.81(2H, dd, J=1.6, 4.3Hz), 8.70(2H, dd, J=1.3, 4.5Hz)(DMSO-d6)	438
XA 2049	2.67 (3H, s), 3.06 (1H, dd, J = 12.4, 10.8 Hz), 3.25 (3H, m), 3.57 (3H, s), 3.62 (2H, m), 4.12 (1H, dd, J = 10.0, 2.0 Hz), 6.68 (1H, s), 7.59 (2H, d, J = 8.0 Hz), 7.80 (1H, dd, J = 4.8, 1.2 Hz), 8.09 (1H, d, J = 8.0 Hz), 8.71 (1H, dd, J = 4.8, 1.2 Hz) (CDCl3)	430
XA 2050	3.05(1H, m), 3.30-3.48(3H, m), 3.64(3H, s), 4.08-4.22(2H, m), 4.68(1H, m), 5.15(1H, d, J= 12.3 Hz), 5.21(1H, d, J= 12.6 Hz), 6.63(1H, s), 7.21(2H, d, J= 8.4 Hz), 7.28-7.39(7H, m), 7.59(2H, d, J=6.3 Hz), 8.68(2H, d, J=6.3 Hz) (CDCI3)	560
XA 2051	2.88-3.34(6H, m), 3.67(3H, s), 4.56(1H, dd, J= 9.9, 3.3 Hz), 6.62(1H, s), 7.19(2H, d, J= 10.8 Hz), 7.36(2H, d, J= 10.8 Hz), 7.58(2H, dd, J=4.5, 1.5 Hz), 8.67(2H, dd, J=4.5, 1.5 Hz) (CDCI3)	426
XA 2052	3.04(1H, m), 3.29-3.48(3H, m), 3.64(3H, s), 4.10-4.15(2H, m), 4.68(1H, m), 5.15(1H, d, J= 12.3 Hz), 5.21(1H, d, J= 12.6 Hz), 6.63(1H, s), 7.21(2H, d, J= 8.1 Hz), 7.32-7.39(7H, m), 7.59(2H, d, J=6.0 Hz), 8.68(2H, d, J=6.0 Hz) (CDCI3)	560
XA 2053	3.01(1H, m), 3.29-3.41(3H, m), 3.66(3H, s), 4.05-4.13(2H, m), 4.67(1H, m), 6.64(1H, s), 7.23(2H, d, J= 8.4 Hz), 7.41(2H, d, J= 8.4 Hz), 7.60(2H, dd, J=4.5, 1.5 Hz), 8.69(2H, dd, J=4.5, 1.5 Hz) (CDCl3)	527
XA 2054	2.28(3H, s), 3.07(4H, m), 3.59(4H, m), 3.73(3H, s), 5.78(1H, s), 6.70(1H, s), 6.98(1H, m), 7.40(1H, m), 7.60-7.66(2H, m), 7.81(2H, dd, J=1.6, 4.3Hz), 8.72(2H, dd, J=1.3, 4.5Hz)(CDCl3)	445

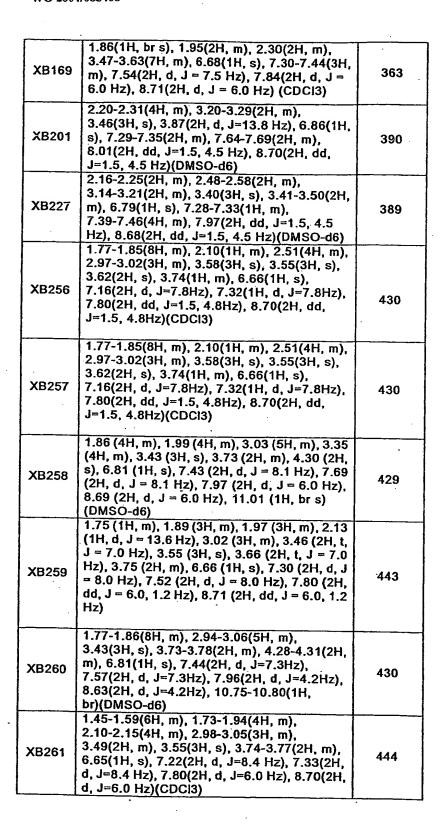




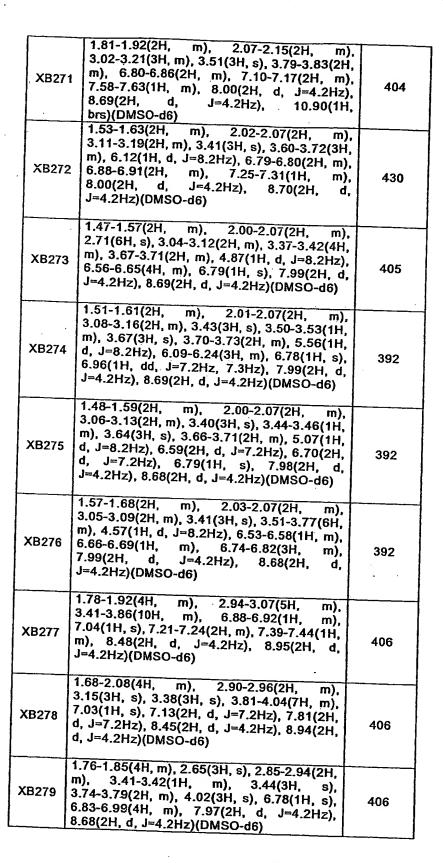
Γ	1,000	
XB48	1.90-2.10(3H, m), 2.32-2.44(1H, m), 3.11-3.20(1H, m), 3.45(1H, dd, J=10.5, 12.6 Hz), 3.57(3H, s), 3.61-3.72(2H, m), 4.08(1H d, J=11.1 Hz), 6.67(1H, s), 7.30-7.35(1H, m), 7.56-7.62(2H, m), 7.74(1H, d, J=13.8 Hz), 7.80(2H, dd, J=1.8, 4.5 Hz), 8.70(2H, dd, J=1.8, 4.8 Hz)(CDCl3)	1 \$
XB49	1.91-2.09(3H, m), 2.37-2.42(1H, m), 3.12-3.19(1H, m), 3.45(1H, dd, J=10.8, 12.9 Hz), 3.57(3H, s), 3.60-3.72(2H, m), 4.08(1H d, J=11.1 Hz), 6.67(1H, s), 7.30-7.35(1H, m) 7.54-7.62(2H, m), 7.75(1H, d, J=8.1 Hz), 7.80(2H, dd, J=1.5, 4.5 Hz), 8.70(2H, dd, J=1.8, 4.5 Hz)(CDCI3)	ł
XB50	1.59-1.67(1H, m), 1.72-1.81(1H, m), 2.08(1H, dt, J=3.4, 12.7 Hz), 2.23-2.40(1H, m), 3.06-3.14(1H, m), 3.41-3.54(2H, m), 3.42(3H, s), 3.93(1H, d, J=14.0 Hz), 7.02(1H, s), 7.24-7.29(1H, m), 7.34-7.39(2H, m), 7.56-7.59(2H, m), 8.55(2H, d, J=6.6 Hz), 8.98(2H, d, J=6.5 Hz)(DMSO-d6)	363
XB80	2.21-2.36(4H, m), 3.19-3.31(2H, m), 3.46(3H, s), 3.88(2H, d, J=13.2 Hz), 6.86(1H, s), 7.38-7.42(1H, m), 7.46-7.51(2H, m), 7.58-7.64(2H, m), 8.01(2H, d, J=5.1 Hz), 8.70(2H, d, J=5.1 Hz)(DMSO-d6)	372
XB122	1.44(2H, m), 1.75-1.83(3H, m), 2.63(2H, d, J = 6.9 Hz), 2.90(2H, m), 3.51(3H, s), 3.64(2H, m), 6.65(1H, s), 7.17-7.34(5H, m), 7.80(2H, d, J = 6.3 Hz), 8.70(2H, d, J = 6.3 Hz) (CDCI3)	361
XB123	1.44-2.16(5H, m), 2.86-2.97(2H, m), 3.49(3H, s), 3.62(1H, m), 3.72(1H, m), 4.48(1H, d, J = 7.2 Hz), 6.64(1H, s), 7.07(2H, m), 7.32(2H, m), 7.79(2H, d, J = 6.3 Hz), 8.69(2H, d, J = 6.3 Hz) (CDCI3)	395
XB124	1.38-1.60(3H, m), 1.78(1H, m), 2.16(1H, m), 2.79-2.94(2H, m), 3.20(3H, s), 3.49(3H, s), 3.59(1H, m), 3.69(1H, m), 3.88(1H, d, J = 7.5 Hz, 1H), 6.64(1H, s), 7.08(2H, m), 7.25(2H, m), 7.79(2H, d, J = 6.0 Hz), 8.70(2H, d, J = 6.0 Hz) (CDC(3)	409
XB127	1.87-2.06(4H, m), 2.79(1H, m), 3.10(2H, m), 3.57(3H, s), 3.78(2H, m), 6.68(1H, s), 7.23-7.29(3H, m), 7.34(2H, m), 7.84(2H, d, J = 6.0 Hz), 8.72(2H, d, J = 6.0 Hz) (CDCI3)	347
XB130	1.61-2.03(4H, m), 2.78(1H, m), 3.09(2H, m), 3.57(3H, s), 3.79(2H, m), 6.69(1H, s), 7.03(2H, m), 7.23(2H, m), 7.84(2H, d, J = 5.4 Hz), 8.72(2H, br s) (CDCl3)	365
XB134	1.78-1.95(4H, m), 2.80-2.91(1H, m), 2.96-3.09(2H, m), 3.45(3H, s), 3.81(2H, d, J=13.1 Hz), 6.80(1H, s), 7.33(1H, dd, J=2.0, 8.3 Hz), 7.56-7.60(2H, m), 7.99(2H, dd, J=1.6, 4.5 Hz), 8.69(2H, dd, J=1.5, 4.5 Hz)(DMSO-d6)	415
	· · · · · · · · · · · · · · · · · · ·	

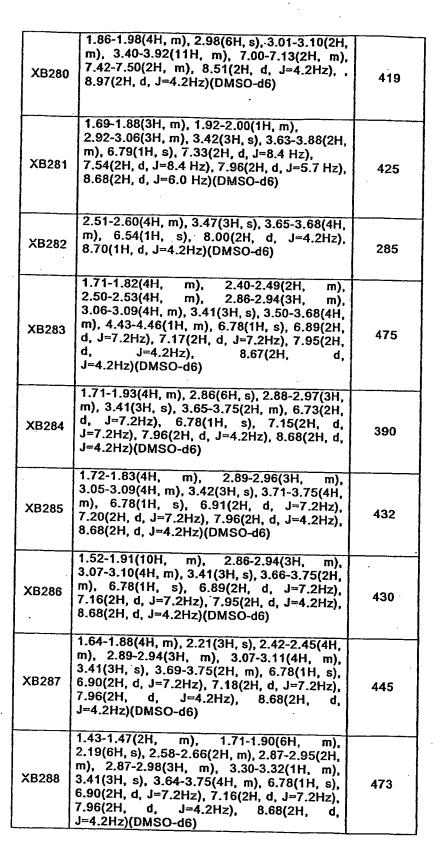


XB145	1.82-2.02(4H, m), 3.09-3.27(3H, m), 3.57(3H, s), 3.79(2H, m), 3.86(3H, s), 6.67(1H, s), 6.89-6.99(2H, m), 7.21-7.26(2H, m), 7.84(2H, d, J = 6.0 Hz), 8.72(2H, d, J = 6.0 Hz) (CDCI3)	377
XB157	1.85-2.07(2H,m), 2.17-2.30(2H,m), 2.91-3.10(1H,m), 3.10-3.24(2H,m), 3.57(3H,s), 3.71-3.88(2H,m), 6.69(1H,s), 6.99-7.06(1H,m), 7.21(1H,dd,J=2.1,8.7Hz), 7.45(1H,s), 7.49-7.65(1H,m), 7.83(2H,dd,J=1.8,4.5Hz), 8.72(2H,dd,J=1.2,4.8Hz)(CDCI3)	405
XB158	2.22-2.32(4H, m), 3.22(2H, m), 3.37(1H, m), 3.58(3H, s), 3.82(2H, m, 6.71(1H, s), 7.10(1H, m), 7.29(1H, m), 7.67(1H, m), 7.83(2H, d, J = 6.3 Hz), 8.72(2H, d, J = 6.3 Hz) (CDCI3)	406
XB159	2.19-2.26(4H, m), 3.21(2H, m), 3.35(1H, m), 3.59(3H, s), 3.82(2H, m), 6.70(1H, s), 6.95(1H, dt, J = 9.0, 2.1 Hz), 7.13(1H, dd, J = 9.0, 2.1 Hz), 7.71(1H, m), 7.85(2H, d, J = 6.3 Hz), 8.72(2H, d, J = 6.3 Hz) (CDCI3)	405
XB160	2.13-2.34(2H,m), 2.34-2.43(2H,m), 3.10-3.38(3H,m), 3.57(3H,s), 3.68-3.83(2H,m), 6.69(1H,s), 7.29-7.40(2H,m), 7.46-7.59(1H,m), 7.64-7.78(1H,m), 7.80-7.78(2H,m), 8.72(2H,d,J=6.0Hz)(CDCI3)	388
XB161 <sub>.</sub>	2.19(2H, m), 2.38(2H, m), 3.18(2H, m), 3.39(1H, m), 3.58(3H, s), 3.80(2H, m), 6.70(1H, s), 7.39(1H, m), 7.50(1H, m), 7.83(2H, d, J = 6.0 Hz), 7.89(1H, d, J = 7.2 Hz), 8.01(1H, d, J = 7.8 Hz), 8.73(2H, d, J = 6.0 Hz) (CDCI3)	404
XB162	1.96(2H, m), 2.88(2H, m), 3.15(2H, m), 3.60(3H, s), 3.85(2H, m), 4.63(1H, m), 6.73(1H, s), 7.13-7.23(3H, m), 7.46(1H, d, J = 7.5 Hz), 7.84(2H, d, J = 6.3 Hz), 8.73(2H, d, J = 6.3 Hz)(CDCl3)	420
XB164	1.64(2H, m), 2.23(2H, m), 3.13(2H, m), 3.50(1H, m), 3.53(3H, s), 3.68(2H, m), 6.58(2H, m), 6.68(1H, s), 6.91(2H, m), 7.81(2H, d, J = 6.0 Hz), 8.72(2H, d, J = 6.0 Hz) (CDCI3)	380
XB165	1.91-1.99(4H, m), 2.84(3H, s), 3.07(2H, m), 3.55(3H, s), 3.77(2H, m), 3.84(1H, m), 6.69(1H, s), 6.75-6.87(3H, m), 7.27(2H, m), 7.82(2H, d, J = 6.3 Hz), 8.72(2H, d, J = 6.3 Hz) (CDCI3)	376
XB168	1.52(2H, m), 1.79(3H, s), 1.96(2H, m), 3.09(2H, m), 3.42(3H, s), 3.64(2H, m), 4.86(1H, m), 6.63(1H, s), 7.09-7.19(4H, m), 7.74(2H, d, J = 6.0 Hz), 8.70(2H, d, J = 6.0 Hz) (CDCl3)	422



XB262	J=8.4 Hz), 7.33(2H, d, J=8.4 Hz), 7.81(2H, d, J=6.0 Hz), 8.71(2H, d, J=6.0 Hz)(CDCl3)	472
XB263	1.77-1.86(4H, m), 2.44(1H, m), 2.80(6H, s), 2.98-3.16(4H, m), 3.42(3H, s), 3.62-3.79(6H, m), 4.42(3H, m), 6.93(1H, s), 7.45(2H, d, J=8.4 Hz), 7.58(2H, d, J=6.0 Hz), 8.82(2H, d, J=6.0 Hz)(DMSO-d6)	i
XB264	0.99(3H, t, J=7.2Hz), 1.20-1.24(6H, m), 1.80-1.93(7H, m), 2.10(1H, m), 2.50-2.55(2H, m), 2.97-3.00(3H, m), 3.55(3H, s), 3.60(2H, s), 3.69-3.74(2H, m), 6.65(1H, s), 7.18(2H, d, J=8.4 Hz), 7.34(2H, d, J=8.4 Hz), 7.80(2H, d, J=6.0 Hz), 8.70(2H, d, J=6.0 Hz)(CDCl3)	486
XB265	1.02(6H, d, J=6.6Hz), 1.23-1.28(5H, m), 1.72-2.15(9H, m), 2.51(1H, m), 2.97-3.08(4H, m), 3.55(3H, s), 3.70(2H, s), 3.74-3.78(2H, m), 6.65(1H, s), 7.18(2H, d, J=7.8 Hz), 7.34(2H, d, J=6.0 Hz), 8.70(2H, d, J=6.0 Hz)(CDCI3)	500
XB266	1.//-1.87(4H, m), 2.44(1H, m), 2.80(6H, s), 2.99-3.09(4H, m), 3.42(3H, s), 3.62-3.79(6H, m), 4.42(3H, m), 6.95(1H, s), 7.45(2H, d, J=8.1 Hz), 7.58(2H, d, J=8.1 Hz), 8.29(2H, d, J=6.0 Hz), 8.86(2H, d, J=6.0 Hz)(DMSO-d6)	473
XB267	1.85-1.88(4H, m), 2.81(1H, m), 2.99-3.07(2H, m), 3.44(3H, s), 3.79-3.84(2H, m), 6.82(1H, s), 7.29(2H, d, J=8.4 Hz), 7.51(2H, d, J=8.4 Hz), 8.01(2H, d, J=6.0 Hz), 8.70(2H, d, J=6.0 Hz)(DMSO-d6)	425
XB268	1.83-1.99(4H, m), 2.83(1H, m), 2.98-3.06(2H, m), 3.45(3H, s), 3.79-3.84(2H, m), 6.82(1H, s), 7.29-7.43(3H, m), 7.53(1H, s), 8.01(2H, d, J=6.0 Hz), 8.70(2H, d, J=6.0 Hz)(DMSO-d6)	425
XB269	1.74-1.96(8H, m), 2.51(1H, m), 2.65-3.01(2H, m), 3.04-3.18(4H, m), 3.44(3H, s), 3.77-3.81(2H, m), 6.49(2H, d, J=8.4 Hz), 6.80(1H, s), 7.09(2H, d, J=8.4 Hz), 8.00(2H, dd, J=4.5, 1.8 Hz), 8.69(2H, dd, J=4.5, 1.8 Hz)(DMSO-d6)	416
XB270	1.83-1.99(8H, m), 2.72(1H, m), 2.97-3.07(2H, m), 3.19-3.23(4H, m), 3.45(3H, s), 3.78-3.83(2H, m), 6.38(1H, d, J=7.8 Hz) 6.44(1H, s), 6.53(1H, d, J=7.5 Hz), 6.81(1H, s), 7.09(1H, dd, J=7.8, 7.8 Hz), 8.00(2H, d, J=5.4 Hz), 8.70(2H, d, J=5.7 Hz)(DMSO-d6)	416









XB289	1.72-1.94(4H, m), 2.92-2.99(3H, m), 3.08-3.11(4H, m), 3.41(3H, s), 3.52-3.56(4H, m), 3.66-3.75(2H, m), 5.11(2H, s), 6.78(1H, s), 6.93(2H, d, J=7.2Hz), 7.20(2H, d, J=7.2Hz), 7.28-7.39(5H, m), 7.95(2H, d, J=4.2Hz), 8.69(2H, d, J=4.2Hz)(DMSO-d6)	565
XB290	1.53-1.63(2H, m), 1.85-1.89(2H, m), 2.14(3H, s), 2.31-2.46(8H, m), 2.86-2.94(2H, m), 3.34-3.35(1H, m), 3.39(3H, s), 3.70-3.74(2H, m), 6.79(1H, s), 7.98(2H, d, J=4.2Hz), 8.69(2H, d, J=4.2Hz)(DMSO-d6)	369
XB291	1.52-1.63(2H, m), 1.85-1.90(2H, m), 2.34-2.42(11H, m), 2.86-2.94(2H, m), 3.39(3H, s), 3.45-3.50(2H, m), 3.70-3.74(2H, m), 4.38-4.40(1H, m), 6.80(1H, s), 7.98(2H, d, J=4.2Hz), 8.69(2H, d, J=4.2Hz)	399
XB292	1.71-1.83(4H, m), 2.81-3.00(11H, m), 3.28-3.30(1H, m), 3.41(3H, s), 3.66-3.75(2H, m), 6.78(1H, s), 6.89(2H, d, J=7.2Hz), 7.17(2H, d, J=7.2Hz), 7.96(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	431
XB293	1.43-1.53(2H, m), 1.93-1.98(3H, m), 2.63-2.66(1H, m), 2.92-3.00(2H, m), 3.39(3H, s), 3.62-3.79(7H, m), 6.78(1H, s), 6.88-6.97(2H, m), 7.18-7.22(1H, m), 7.35(1H, d, J=7.3Hz), 7.98(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	406
XB294	1.42-1.53(2H, m), 1.96-2.08(3H, m), 2.61-2.67(1H, m), 2.91-2.99(2H, m), 3.39(3H, s), 3.62-3.80(7H, m), 6.77(1H, s), 6.86(2H, d, J=7.2Hz), 7.25(2H, d, J=7.2Hz), 7.97(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	406
XB295	1.81-1.91(2H, m), 2.61-2.20(2H, m), 2.96-3.17(6H, m), 3.41-3.47(5H, m), 3.74-3.86(4H, m), 6.90-7.03(3H, m), 7.21-7.29(2H, m), 8.44(2H, d, J=4.2Hz), 8.93(2H, d, J=4.2Hz), 9.30-9.38(2H, br)(DMSO-d6)	420
XB296	1.80-1.91(2H, m), 2.07-2.21(2H, m), 2.96-3.11(6H, m), 3.34-3.41(5H, m), 3.69-3.86(4H, m), 6.91(2H, d, J=7.2Hz), 7.05(1H, s), 7.20(2H, d, J=7.2Hz), 8.49(2H, d, J=4.2Hz), 8.96(2H, d, J=4.2Hz), 9.44-9.50(2H, br)(DMSO-d6)	420



XB297	1.41-1.51(2H, m), 1.91-1.96(3H, m), 2.61-2.65(1H, m), 2.86(6H, s), 2.91-2.98(2H, m), 3.38(3H, s), 3.61-3.67(4H, m), 6.70(2H, d, J=7.2Hz), 6.77(1H, s), 7.20(2H, d, J=7.2Hz), 7.97(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	419
XB298	2.04(2H, d, J=13.1Hz), 2.34(3H, s), 2.53(2H, m), 2.91(2H, m), 3.55(3H, s), 3.70(2H, d, J=13.1Hz), 4.27(1H, m), 6.08(1H, s), 6.86(1H, s), 7.36-7.48(5H, m), 7.80(2H, dd, J=1.6, 4.3Hz), 8.69(2H, dd, J=1.3, 4.5Hz)(CDCl3)	426
XB299	2.06(2H, d, J=13.1Hz), 2.22(2H, m), 2.99(2H, m), 3.13(1H, m), 3.54(3H, s), 3.70(2H, d, J=13.1Hz), 6.68(1H, s), 7.25(1H, s), 7.44-7.48(2H, m), 7.64-7.67(3H, m), 7.78(2H, dd, J=1.6, 4.3Hz), 8.69(2H, dd, J=1.3, 4.5Hz)(CDCI3)	413
XB300	1.75-1.85(4H, m), 2.97-3.10(5H, m), 3.43(3H, s), 3.71-3.76(2H, m), 3.88-3.93(2H, m), 6.70(1H, dd, J=7.2, 7.3Hz), 6.79(1H, s), 7.02-7.06(2H, m), 7.15-7.23(3H, m), 7.31-7.35(2H, m), 7.97(2H, d, J=4.2Hz), 8.69(2H, d, J=4.2Hz)(DMSO-d6)	464
XB301	1.09-1.34(5H, m), 1.57-1.88(9H, m), 2.78-2.93(3H, m), 3.08-3.18(1H, m), 3.41(3H, s), 3.62-3.74(2H, m), 5.27(1H, d, J=8.2Hz), 6.52(2H, d, J=7.2Hz), 6.79(1H, s), 7.01(2H, d, J=7.2Hz), 7.96(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	444
XB302	1.10-1.16(1H, m), 1.32-1.46(4H, m), 1.64-1.82(9H, m), 2.68(3H, s), 2.82-2.93(3H, m), 3.41(3H, s), 3.54-3.74(3H, m), 6.72(2H, d, J=7.2Hz), 6.78(1H, s), 7.12(2H, d, J=7.2Hz), 7.95(2H, d, J=4.2Hz), 8.68(2H, d, J=4.2Hz)(DMSO-d6)	458

No.	NMR	MS[M+1]
YA0262	(DMSO-d6): 3.47(3H, s), 3.48-3.66(4H, m), 3.89-4.02(2H, m), 4.98(1H, m), 7.06(1H, s), 7.35-7.59(3H, m), 7.99(1H, dd, J=7.2, 6.9Hz), 8.25(1H, dd, J=5.4, 1.2Hz), 9.01(1H, d, J=5.1Hz), 9.31(1H, s), 9.84(1H, m), 10.19(1H, m).	367
YA0263	(CDCl3):3.01(1H,dd,J=10.5,12.4Hz), 3.10-3.35(3H,m), 3.57(3H,s), 3.55-3.65(2H,m), 4.05(1H,dd,J=2.4,10.4Hz), 7.00-7.10(1H,m), 7.30(1H,s), 7.22(2H,m), 7.30-7.42(2H,m), 8.15(1H,dd,J=1.3,5.2Hz), 8.86(1H,d,J=5.2Hz), 9.27(1H,d,J=1.0Hz).	367



	<b></b>	
	2.83(1H, dd, J=11.0, 11.9 Hz), 2.93(1H, s),	
	2.99-3.10(3H, m), 3.45(3H, s), 3.61-3.69(2H, m),	
	3.95(1H, dd, J=2.1, 10.3 Hz), 6.97(1H, s),	
YA0264	7.19(2H, t, J=8.8 Hz), 7.48-7.56(2H, m), 8.17(1H,	367
	dd, J=1.0, 5.0 Hz), 8.99(1H, d, J=5.1 Hz),	
	9.29(1H, d, J=1.0 Hz)(DMSO-d6)	
•	, , , , , , , , , , , , , , , , , , , ,	
	3.39-3.47(2H, m), 3.45(3H, s), 3.55-3.66(2H, m),	
	3.86-3.96(2H, m), 4.64-4.71(1H, m), 7.05(1H, s),	
YA0264	7.36(2H, t, J=8.7 Hz), 7.77-7.81(2H, m), 8.23(1H,	
(HCI)	dd, J=1.2, 5.1 Hz), 9.02(1H, d, J=5.1 Hz),	367
(,	9.32(1H, d, J=1.2 Hz), 9.79(1H, d, J=10.2 Hz),	
	10.13-10.28(1H, m)(DMSO-d6)	
	(CDCl3) :2.81(1H,dd,J=10.5,12.6Hz),	
	3.15-3.40(3H,m),	
	3.50-3.65(4H,m),3.65-3.80(1H,m),	
YA0267	4.51(1H,dd,J=2.7,10.5Hz), 7.20-7.45(4H,m),	383
	7.74(1H,dd,J=1.5,7.5Hz), 8.15-8.20(1H,m),	
	8.85(1H,d,J=5.1Hz), 9.27(1H,s).	
	(CDCI3) :3.00(1H,dd,J=10.5,12.6Hz),	
YA0268	3.10-3.35(3H,m), 3.50-3.70(5H,m),	202
1 AU200	4.03(1H,dd,J=2.4,10.5Hz), 7.32(4H,m),	383
	7.50(1H,s), 8.15(1H,dd,J=1.2,5.1Hz),	
	8.87(1H,d,J=5.1Hz), 9.27(1H,d,J=1.5Hz).	
	3.40-3.50(2H, m), 3.45(3H, s), 3.53-3.65(2H, m),	
	3.87-3.97(2H, m), 4.68(1H, t, J=10.2 Hz),	•
YA0269	7.05(1H, s), 7.59(2H, d, J=11.1 Hz), 7.75(2H, d,	383
	J=11.1 Hz), 8.22(1H, dd, J=1.5, 5.4 Hz), 9.02(1H,	
	d, J=5.1 Hz), 9.31(1H, s), 9.83(1H, d, J=9.6 Hz),	
<del></del>	10.11-10.25(1H, m)(DMSO-d6)	
	(DMSO-d6):3.45(3H,s), 3.40-3.70(4H,m),	
	3.92(2H,t,J=14.1Hz), 4.67(1H,br s), 7.06(1H,s),	
YA0274	7.68(2H,d,J=10.0Hz), 7.72(2H,d,J=10.0Hz),	427
	8.22(1H,d,J=4.8Hz), 9.03(1H,d,J=4.8Hz),	
	9.31(1H,s), 9.88(1H,br s), 10.22(1H,br s).	
	3.38-3.57(4H, m), 3.35(3H,s), 3.89(3H,s),	
	3.91-3.97(2H, m), 4.84-4.94(1H, m), 7.06(1H, s),	
YA0289	7.08-7.15(1H, m), 7.18(1H, d, J=8.4 Hz),	379
	7.41-7.49(1H, m), 7.68(1H, d, J=7.6 Hz),	0.0
	8.25(1H, d, J=4.9 Hz), 9.04(1H, d, J=5.1 Hz),	
	9.32(1H, s)(DMSO)	
	(DMSO-d6) :3.40-3.75(7H,m),	
	3.92(2H,t,J=13.2Hz), 4.64(1H,t,J=9.1Hz),	
	7.00-7.10(2H,m), 7.23(1H,d,J=7.6Hz),	
YA0290	7.35(1H,s), 7.42(1H,t, J=7.8Hz),	379
	8.23(1H,d,J=5.6Hz), 9.02(1H,d,J=5.2Hz),	
	9.32(1H,s), 9.65-9.80(1H,brd),	
	9.90-10.15(1H,brd).	
	(DMSO-d6): 3.42(3H, s), 3.36-3.58(4H, m),	
	3.79(3H, s), 3.83-3.95(2H, m), 4.61(1H, m),	
YA0291	7.05(1H, s), 7.07(2H, d, J=8.1Hz), 7.60(2H, d,	379
	J=8.7Hz), 8.22(1H, dd, J=5.1, 1.2Hz), 9.02(1H,	
	d, J=5.4Hz), 9.31(1H, s), 9.58-9.74(2H, m).	
	14,0 0.7112, 0.01(111, 3/, 3.00-3.17(211, 111).	L

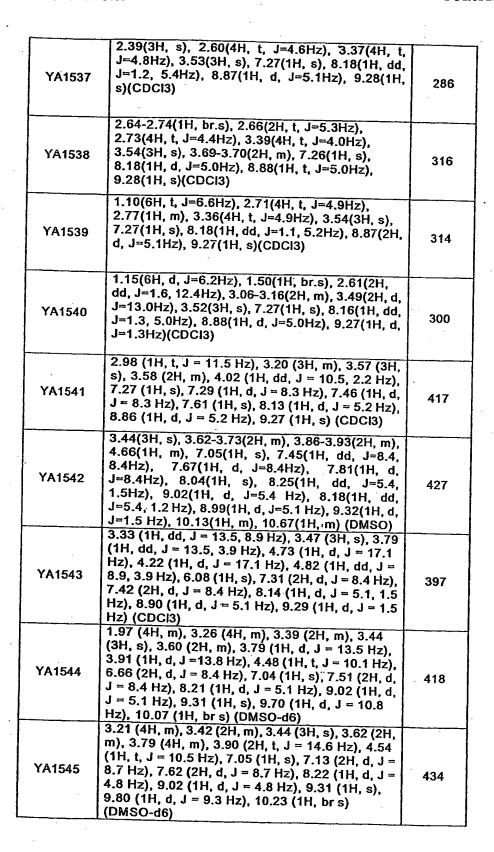
	· · · · · · · · · · · · · · · · · · ·	
	1.31(3H, t, J=6.8 Hz), 3.44-3.59(2H, m), 3.48(3H,	
YA0294	s), 3.87-3.97(2H, m), 4.09-4.20(2H, m),	
	4.80-4.91(1H, m), 7.06(1H, s), 7.09-7.17(2H, m),	
	7.44(1H, t, J=7.4 Hz), 7.64(1H, d, J=7.5 Hz),	393
	8 23/1H d l=5 2 H=\ 0.02/4H d l=5 2 H=\	
	8.23(1H, d, J=5.3 Hz), 9.03(1H, d, J=5.2 Hz),	
	9.32(1H, s), 9.49-9.60(2H, m)(DMSO-d6)	
	(DMSO-d6):3.45(3H,s), 3.64(3H,m), 3.93(3H,m),	
	4.78(1H,t,J=9.6Hz), 7.13(1H,s),	
V40004	7.97(2H,d,J=8.7Hz), 8.01(2H,d,J=8.7Hz),	
YA0304	8.43(2H,d,J=6.2Hz), 8.93(2H,d,J=6.2Hz),	374
	10.12(1H,s), 10.70(1H,s).	
	10.12(11,15), 10.10(111,15).	
	(CDCl3):2.00(4H,m), 3.05(1H,t,J=11.7Hz),	
	3.18-3.30(3H,m), 3.29(4H,m), 3.56(3H,s),	•
)/a aaa 4	3.62(2H,m), 3.91(1H,d,J=8.4Hz),	
YA0331	6.57(2H,d,J=8.7Hz), 7.31(3H,m),	418
	8.17(1H,dd,J=1.2,5.1Hz), 8.85(1H,d,J=5.1Hz),	
	9.27(1H,d,J=1.2Hz).	
	(CDCl3):3.02(1H,dd,J=10.8,12.6Hz),	
	3.18(8H,m), 3.56(3H,s), 3.61(1H,t,J=9.0Hz),	
V40007	3.87(4H,m), 3.95(1H,dd,J=2.7,10.8Hz),	
YA0337	6.93(2H,d,J=8.9Hz), 7.31(1H,s),	434
	7.36(2H,d,J=8.9Hz), 8.16(1H,dd,J=1.5,5.4Hz),	
	8.85(1H,d,J=5.4Hz), 9.27(1H,d,J=1.5Hz).	
	(CDCl3):2.36(3H,s), 2.59(4H,m),	•
•	3.02(1H,t,J=11.4Hz), 3.16-3.29(7H,m),	
YA0340	3.26(3H,s), 3.61(2H,m), 3.94(1H,d,J=8.0Hz),	447
1710010	6.94(2H,d,J=8.7Hz), 7.31(1H,s),	441
	7.34(2H,d,J=8.7Hz), 8.16(1H,d,J=5.1Hz),	
•	8.85(1H,d,J=5.1Hz), 9.27(1H,s).	
	3.39-3.50(2H, m), 3.47(3H, s), 3.61-3.73(1H, m),	· · · · · · · · · · · · · · · · · · ·
	3.78(3H, s), 3.83(3H, s), 3.87-3.92(3H, m),	
	4.92(1H, t, J=10.5 Hz), 6.99-7.11(3H, m),	
YA0361		400
170301	7.57(1H, d, J=2.7 Hz), 8.25(1H, dd, J=1.2, 5.1	409
	Hz), 9.03(1H, d, J=4.8 Hz), 9.31(1H, d, J=0.9	
	Hz), 9.78(1H, d, J=9.0 Hz), 10.21-10.38(1H,	
	m)(DMSO-d6)	
	(DMSO-d6): 3.47(3H, s), 3.37-4.04(6H, m),	
	3.94(6H, s), 5.09(1H, m), 6.82(2H, d, J=8.4Hz),	
YA0362	7.05(1H, s), 7.45(1H, t, J=8.4Hz), 8.22(1H, m),	409
	8.24(1H, dd, J=5.4, 1.5Hz), 9.05(1H, d,	
	J=5.1Hz), 9.32(1H, s), 10.06(1H, m).	
	3.38-3.60(4H, m), 3.47(3H, s), 3.88-3.95(2H, m),	
	3.90(3H, s), 4.86-4.92(1H, m), 6.96-7.01(1H, m),	
YA0366	7 06(4U a) 7 42(4U d l=0 0 U=) 7 74 7 70(4U	207
170000	7.06(1H, s), 7.12(1H, d, J=8.8 Hz), 7:71-7.79(1H,	397
	m), 8.23-8.24(1H, m), 9.03(1H, d, J=5.1 Hz),	
	9.32(1H, d, J=1.2 Hz), 9.55-9.72(2H, m)(DMSO)	<u> </u>
	(DMSO-d6):3.30-3.75(7H,m), 3.80-4.00(5H,m),	
	4.80-5.00(1H,m), 6.93-7.00(1H,m), 7.05(1H,s),	
YA0367/	7.11(1H,dd,J=2.4,11.4Hz), 7.84(1H,m),	207
YA0368	8.23(1H,d,J=5.1Hz), 9.03(1H,d,J=5.1Hz),	397
	9.31(1H,s), 9.60-9.80(1H,brd),	
	9.90-10.15(1H,brd).	



	3.31-3.56(3H, m), 3.45(3H, s), 3.69-3.78(1H, m),	
.YA0370	3.90-3.99(2H, m), 3.94(3H, s), 4.95-5.03(1H, m),	
	6.96-7.02(1H, m), 7.03-7.09(2H, m),	
	7.49-7.56(1H, m), 8.24(1H, d, J=4.4 Hz).	397
	8.51-8.69(1H, m), 9.03(1H, d, J=5.1 Hz),	
L_	9.32(1H, s), 10.55-10.67(1H, m) (DMSO)	
	2.77(1H, dd, J=10.5, 12.0 Hz), 3.18-3.30(3H, m),	
1	3.61(3H, s), 3.64-3.71(2H, m), 3.86(3H, s),	
V40070	4.37(1H, dd, J=2.1, 10.1 Hz), 6.89(1H, d, J=1.7	
YA0378	Hz), 6.99(1H, dd, J=1.6, 8.2 Hz), 7.32(1H, s),	413
	7.50(1H, d, J=8.2 Hz), 8.19(1H, d, J=5.2 Hz),	
	8.86(1H, d, J=5.2 Hz), 9.27(1H, s)(CDCl3)	
	(CDCl3) :2.76(1H,dd,J=10.2,12.3Hz),	
	3.10-3.40(3H,m), 3.55-3.80(5H,m), 3.85(3H,s),	
	4.39(1H,dd,J=2.4,10.2Hz), 6.78(1H,d,J=8.7Hz),	
YA0399	7.32(1H,s), 7.39(1H,dd,J=2.7,8.7Hz),	457
	7.72(1H,d,J=2.4Hz), 8.20(1H,dd,J=1.2,5.1Hz),	
l	8.87(1H,d,J=5.1Hz), 9.27(1H,d,J=1.2Hz).	:
	(CDCl3): 1.98-2.03(4H, m), 2.84(1H, m),	
1	3.17-3.32(7H, m), 3.60(3H, s), 3.59-3.71(2H, m),	
	3.85(3H, s), 4.28(1H, d, 8.4Hz), 6.10(1H, d,	
YA0408	1=1 8H=1 6 19(1H d 1=0 2H=1 7 20(4H =1	448
	J=1.8Hz), 6.18(1H, d, J=8.3Hz), 7.29(1H, s),	
	7.33(1H, d, J=8.4Hz), 8.21(1H, d, J=5.2Hz),	
	8.85(1H, d, J=5.2Hz), 9.27(1H, s).	<del></del>
	(CDCl3):1.95-2.10(4H,m), 2.95-3.10(1H,m),	
	3.19-3.45(7H,m), 3.59(3H,s), 3.50-3.80(2H,m),	
YA0409	3.80(3H,s), 4.48(1H,dd,J=2.2,10.2Hz),	448
İ	6.49(1H,dd,J=3.0,8.9Hz), 6.63-6.87(2H,m),	
	7.32(1H,s), 8.20(1H,dd,J=1.4,5.2Hz), 8.86(2H,d,J=5.2Hz), 9.27(1H,d,J=1.1Hz).	
	(CDCl3):3.14(2H,m), 3.22(1H,t,J=11.6Hz),	
	3.41(1H,t,J=11.6Hz), 3.82(2H,m), 3.83(3H,s),	
j	3.88(3H,s), 4.58(1H,dd,J=3.1,11.0Hz),	·
YA0414	6.51(2H,m), 7.32(1H,s),	415
	8 10(1H dd (-4 5 5 2H-) 9 00(4H d (-5 0H-)	
	8.19(1H,dd,J=1.5,5.3Hz), 8.86(1H,d,J=5.3Hz), 9.27(1H,d,J=1.5Hz).	
	(DMSO-d6):3.35-3.70(4H,m), 3.48(3H,s),	•
	3.78(3H,s), 3.97(2H,m), 4.70(1H,m),	
YA0423	7.06(1H,t,J=7.7Hz), 7.07(1H,s),	
- 17(0423	7.15(1H,d,J=7.7Hz), 7.31(1H,d,J=7.7Hz),	455
	7.39(1H,t,J=7.7Hz), 7.61(2H,d,J=8.1Hz),	1
	7.70(2H,d,J=8.1Hz), 8.25(1H,d,J=4.5Hz),	
	9.07(1H,d,J=4.5Hz), 9.33(1H,s), 9.66(1H,br s).	
	(DMSO-d6):3.61(3H,m), 3.76(3H,s), 3.81(3H,s),	•
•	4.01(3H,m), 4.69(1H,t,J=9.9Hz),	
YA0425	7.05(2H,d,J=9.0Hz), 7.07(1H,s),	
170425	7.67(2H,d,J=9.0Hz), 7.76(4H,s),	455
	8.24(1H,dd,J=1.2,5.1Hz), 9.03(1H,d,J=5.1Hz),	l
	9.32(1H,d,J=1.2Hz), 9.79(1H,d,J=10.2Hz),	ł
	10.07(1H,s).	
	(DMSO-d6):3.30-3.70(4H,m), 3.42(3H,s),	1
YA0434	3.96(2H,d,J=13.8Hz), 4.71(1H,t,J=11.3Hz),	
1 70434	7.06(1H,s), 7.33(2H,t,J=8.0Hz), 7.77(6H,m),	443
	8.24(1H,d,J=5.4Hz), 9.03(1H,d,J=5.4Hz),	1
	9.32(1H,s), 9.80(1H,d,J=8.7Hz), 10.03(1H,s).	

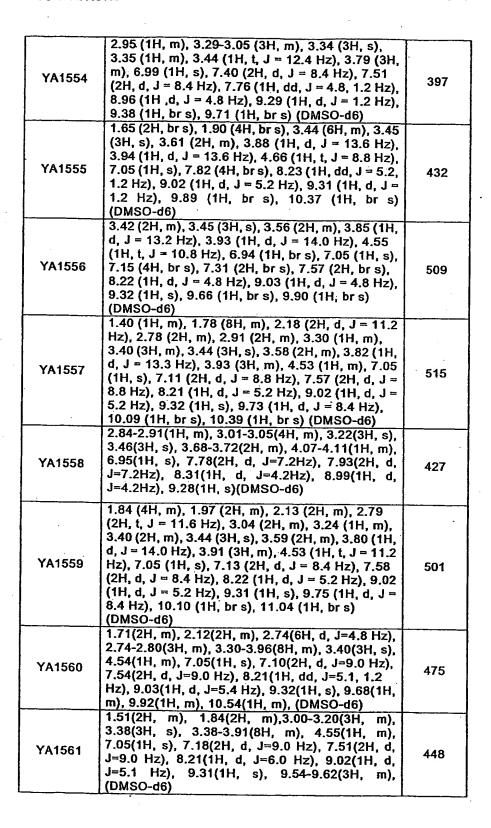
	3.43-3.59(2H, m), 3.48(3H, s), 3.63-3.75(2H, m),	
YA0442	3.97-4.01(2H, m), 4.80-4.86(1H, m), 7.06(1H, s),	• *
	7.60-7.64(2H, m), 7.86-7.88(1H, m),	
	7.95-8.00(2H, m), 8.05-8.07(1H, m),	399
	8.24-8.27(2H, m), 9.02(1H, d, J=5.4 Hz),	
	9.32(1H, s), 10.01(1H, d, J=10.2 Hz),	
	10.30-10.41(1H, m)(DMSO-d6)	
	(CDCI3): 2.97(1H, dd, J=12.3, 10.5Hz),	
	3.18-3.28(5H, m), 3.58(3H, s), 3.59(1H, m),	
YA0517	3.77(1H, m), 4.27(1H, dd, 10.2, 2.7Hz),	391
1 AUS 17	4.62(2H, m), 6.89(1H, t, J=7.5Hz), 7.16(1H, m),	331
	7.27(1H, m), 7.28(1H, s), 8.26(1H, dd, J=5.4,	
	1.5Hz), 8.86(1H, d, J=5.4Hz), 9.26(1H, s).	
	(DMSO-d6):3.15-3.35(1H,m), 3.38-3.50(4H,m),	
	3.70-4.30(9H,m), 5.00-5.20(1H,m),	
YA0864	7.00-7.10(2H,m), 7.10-7.20(1H,m),	487
	7.30-7.50(6H,m), 8.15-8.20(1H,m),	
	8.30-8.40(1H,brd), 9.05(1H,d,J=5.1Hz),	
	9.31(1H,d,J=0.9Hz).	
	(CDCl3):1.80-2.40(3H, m), 3.12-3.34(4H, m),	
	3.39-4.20(7.6H, m), 4.50-5.07(0.6H, m),	
	5.30-5.60(0.7H, m), 5.72-6.05(0.1H, m),	
YA1074	6.52-6.80(2H, m), 6.82-7.22(1H, m), 7.28(1H, s),	439
	8.18(1H, d,J=4.8Hz), 8.89(1H, d,J=5.1Hz),	
	9.28(1H, d,J=1.2Hz)	
	(CDCl3):2.50-2.62(1H,m), 2.80-2.95(1H,m),	
	3.02-3.20(1H,m), 3.25-3.40(1H,m),	
YA1339	3.50-3.74(5H,m), 3.75-3.80(1H,m), 3.85(3H,s),	411
INIOOS	6.60-6.80(2H,m), 7.30(1H,s),	711
	7.48(1H,t,J=8.4Hz), 8.19(1H,dd,J=1.2,5.1Hz),	
	8.86(1H,d,J=5.1Hz), 9.27(1H,d,J=1.5Hz).	
	(DMSO-d6) :2.55(3H,d,J=3.9Hz),	
	3.40-3.80(3H,m), 3.45(3H,s), 3.80-4.15(6H,m),	
YA1340/	4 05 5 45(4U m) 6 00 7 05(4U m) 7 05(4U c)	
	4.85-5.15(1H,m), 6.90-7.05(1H,m), 7.05(1H,s),	411
YA1341	7.13(1H,dd,J=2.4,11.4Hz),	
	8.21(1H,dd,J=1.2,5.1Hz), 9.04(1H,d,J=5.1Hz),	
	9.31(1H,d,J=1.2Hz), 11.50-12.20(1H,brd).	
	2.90-3.10 (1H, m), 3.15-3.35 (3H, m), 3.50-3.70	
•	(5H, m), 3.80-4.05 (7H, m), 6.87 (1H, d, J = 8.1	
YA1534	Hz), 6.90-7.10 (2H, m), 7.31 (1H, s), 8.16 (1H, d,	408
TA 1034	J = 4.6 Hz), 8.85 (1H, d, J = 5.0 Hz), 9.27 (1H, s)	400
	(CDCI3)	
	(0000)	
	3.45 (3H, s), 3.46 (2H, m), 3.64 (m, 2H), 3.91 (2H,	
	t, J = 16.1 Hz), 4.68 (1H, t, J = 10.5 Hz), 7.05 (1H,	
	s), 7.59 (2H, d, J = 8.4 Hz), 7.79 (2H, d, J = 8.4	
YA1535	Hz), 8.23 (1H, dd, J = 5.1, 1.2 Hz), 9.02 (1H, d, J	383
•		
	= 5.1 Hz), 9.31 (1H, d, J = 1.2 Hz), 10.00 (1H, d,	
	J = 8.7 Hz), 10.49 (1H, br s) (DMSO-6)	
	3.45 (3H, s), 3.46 (2H, m), 3.64 (m, 2H), 3.91 (2H,	
	t, $J = 16.1 Hz$ ), $4.68 (1H, t, J = 10.5 Hz), 7.05 (1H, t)$	
VALEGO	s), $7.59$ (2H, d, $J = 8.4$ Hz), $7.79$ (2H, d, $J = 8.4$	200
YA1536	Hz), $8.23$ (1H, dd, $J = 5.1$ , $1.2$ Hz), $9.02$ (1H, d, $J$	383
	= 5.1  Hz), 9.31 (1H, d, J = 1.2 Hz), 10.00 (1H, d,	
	J = 8.7 Hz), 10.49 (1H, br s) (DMSO-6)	1

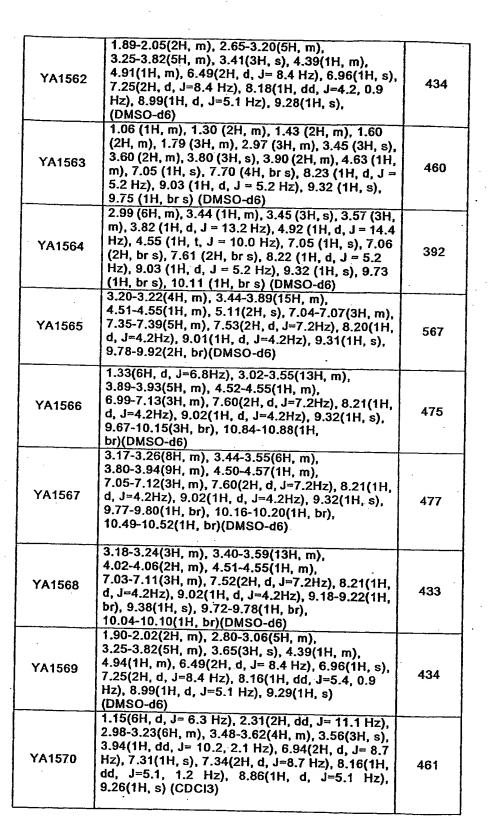
### WO 2004/085408





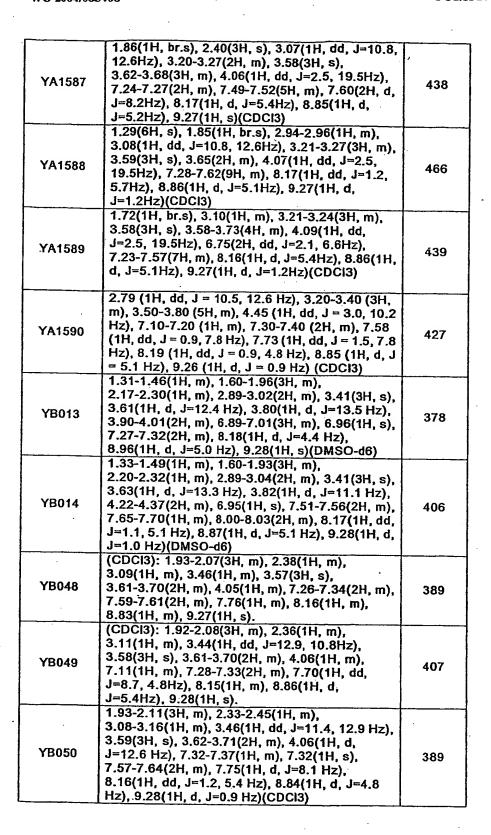
	T 20 (0)	
YA1546	2.80 (3H, d, J = 4.5 Hz), 3.26 (4H, m), 3.44 (3H, s), 3.45 (4H, m), 3.60 (2H, m), 3.80 (1H, d, J = 3.5 Hz), 3.90 (3H, m), 4.54 (1H, t, J = 10.5 Hz), 7.04 (1H, s), 7.10 (2H, d, J = 8.7 Hz), 7.62 (2H, d, J = 8.7 Hz), 8.20 (1H, dd, J = 5.1, 1.2 Hz), 9.02 (1H, d, J = 5.1 Hz), 9.32 (1H, d, J = 1.2 Hz), 9.86 (1H, d, J = 10.2 Hz), 10.33 (1H, br s), 11.15 (1H, br s) (DMSO-d6)	447
YA1547	2.28(3H, s), 3.07(4H, t, J=4.7Hz), 3.37(4H, t, J=4.8Hz), 3.75(3H, s), 5.76(1H, s), 7.26-7.33(2H, m), 7.45(2H, dd, J=7.8, 7.8Hz), 7.79(2H, d, J=7.8Hz), 8.14(1H, d, J=5.4Hz), 8.87(1H, dd, J=7.8, 7.8Hz), 9.28(1H, d, J=1.2Hz)(CDCI3)	428
YA1548	2.37 (1H, m), 2.43 (1H, m), 2.80 (3H, d, J = 5.2 Hz), 2.81 (3H, d, J = 5.2 Hz), 3.28 (1H, q, J = 8.8 Hz), 3.40 (2H, m), 3.44 (3H, s), 3.57 (5H, m), 3.79 (1H, d, J = 11.4 Hz), 3.97 (2H, m), 4.50 (1H, t, J = 10.0 Hz), 6.69 (2H, d, J = 8.4 Hz), 7.05 (1H, s), 7.54 (2H, d, J = 8.4 Hz), 8.20 (1H, dd, J = 4.8, 1.2 Hz), 9.03 (1H, d, J = 4.8 Hz), 9.32 (1H, d, J = 1.2 Hz), 9.71 (1H, br s), 10.06 (1H, br s), 11.35 (1H, br s) (DMSO-d6)	461
YA1549	2.33 (1H, m), 2.41 (1H, m), 2.79 (3H, d, J = 4.8 Hz), 2.81 (3H, d, J = 4.8 Hz), 3.28 (1H, d, J = 8.4 Hz), 3.39 (2H, m), 3.44 (3H, s), 3.57 (5H, m), 3.79 (1H, d, J = 13.3 Hz), 3.97 (2H, m), 4.50 (1H, t, J = 11.6 Hz), 6.69 (2H, d, J = 8.4 Hz), 7.04 (1H, s), 7.55 (2H, d, J = 8.4 Hz), 8.21 (2H, d, J = 5.2 Hz), 9.02 (2H, d, J = 5.2 Hz), 9.32 (1H, s), 9.75 (1H, br s), 10.14 (1H, br s), 11.45 (1H, br s) (DMSO-d6)	461
YA1550	3.47 (3H, s), 3.60 (2H, m), 3.76 (2H, m), 3.81 (3H, s), 3.94 (2H, m), 4.68 (1H, m), 7.05 (2H, d, J = 8.6 Hz), 7.06 (1H, s), 7.67 (2H, d, J = 8.6 Hz), 7.76 (4H, s), 8.25 (1H, d, J = 5.0 Hz), 9.03 (1H, d, J = 5.0 Hz), 9.32 (1H, s) (DMSO-d6)	455
YA1551	1.18 (1H, m), 1.40 (4H, m), 1.70 (1H, m), 1.80 (4H, m), 2.55 (1H, m), 3.43 (2H, m), 3.45 (3H, s), 3.60 (2H, m), 3.91 (2H, m), 4.60 (1H, t, J = 10.8 Hz), 7.05 (1H, s), 7.35 (2H, d, J = 8.0 Hz), 7.64 (2H, d, J = 8.0 Hz), 9.03 (1H, d, J = 4.8 Hz), 9.31 (1H, s), 9.80 (1H, d, J = 8.8 Hz), 10.24 (1H, m) (DMSO-d6)	431
YA1552	3.02(4H, m), 3.23(4H, m), 3.49(3H, s), 7.08-7.67(10H, m), 8.15(1H, d, J=5.1Hz), 8.87(1H, d, J=5.1Hz), 9.27(1H, s)(CDCl3)	424
YA1553	2.90 (1H, dd, J = 13.2, 9.6 Hz), 3.16 (2H, m), 3.24 (1H, d, 14.4 Hz), 3.31 (3H, s), 3.34 (1H, d, J = 13.6 Hz), 3.47 (1H, t, J = 13.2 Hz), 3.80 (3H, m), 6.97 (1H, s), 7.38 (2H, m), 7.45 (3H, m), 7.64 (1H, dd, J = 5.2, 1.2 Hz), 8.94 (1H, d, J = 5.2 Hz), 9.28 (1H, d, J = 1.2 Hz), 9.54 (1H, br s), 9.78 (1H, br s) (DMSO-d6)	363



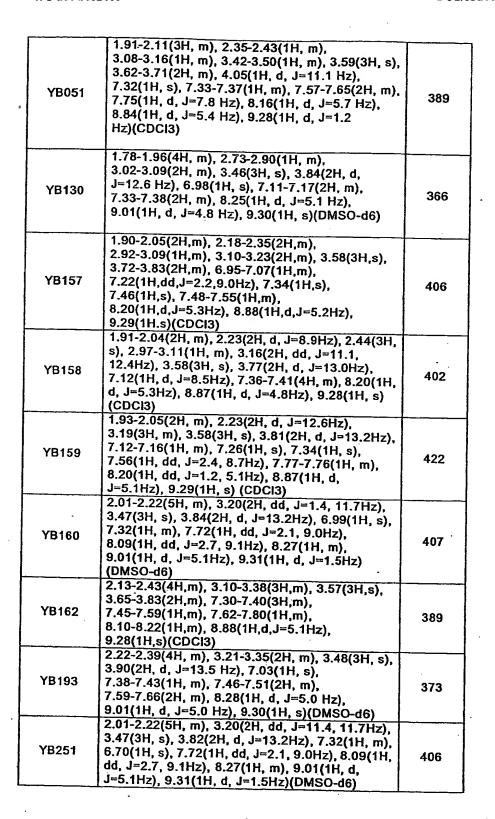


YA1571	1.27(6H, d, J= 6.0 Hz), 2.43(2H, dd, J= 11.1, 11.1 Hz), 3.02(1H, dd, J=12.0, 10.5 Hz), 3.17-3.23(3H, m), 3.45-3.61(4H, m), 3.56(3H, s), 3.81(1H, m), 3.95(1H, m), 6.92(2H, d, J= 8.7 Hz), 7.32(1H, s), 7.35(2H, d, J= 8.7 Hz), 8.17(1H, m), 8.86(1H, d, J=5.1 Hz), 9.26(1H, d, J=1.2 Hz) (CDCl3)	462
YA1572	3.27-3.32(8H, m), 3.47(3H, s), 3.82-3.86(2H, m), 4.36-4.39(1H, m), 7.02(1H, s), 7.72(2H, d, J=7.2Hz), 7.84(2H, d, J=7.2Hz), 7.96-8.04(4H, m), 8.22(1H, d, J=4.2Hz), 9.01(1H, d, J=4.2Hz), 9.30(1H, s)(DMSO-d6)	503
YA1573	2.93-3.10(5H, m), 3.46(3H, s), 3.69-3.71(1H, m), 4.01-4.04(1H, m), 6.99(1H, s), 7.63(2H, d, J=7.2Hz), 7.77(2H, d, J=7.2Hz), 7.88-7.95(4H, m), 8.18(1H, d, J=4.2Hz), 8.99(1H, d, J=4.2Hz), 9.29(1H, s)(DMSO-d6)	450
YA1574	3.08 (1H, dd, J =12.5, 10.4 Hz), 3.24 (3H, m), 3.59 (3H, s), 3.66 (2H, m), 4.09 (1H, dd, J = 10.4, 2.4 Hz), 7.29 (2H, d, J = 8.3 Hz), 7.33 (1H, s), 7.54 (2H, d, J = 8.3 Hz), 7.56 (2H, d, J = 8.3 Hz), 7.59 (2H, d, J = 8.3 Hz), 8.17 (1H, d, J = 4.9 Hz), 8.86 (1H, d, J = 4.9 Hz), 9.27 (1H, s) (CDCI3)	509
YA1575	3.08 (1H, dd, J = 12.4, 10.0 Hz), 3.25 (3H, m), 3.59 (3H, s), 3.67 (2H, m), 4.11 (1H, dd, J = 10.0, 2.0 Hz), 7.33 (1H, s), 7.57 (2H, d, J = 8.0 Hz), 7.63 (2H, d, J = 8.0 Hz), 7.71 (4H, s), 8.16 (1H, dd, J = 5.2, 1.2 Hz), 8.16 (1H, dd, J = 5.2, 1.2 Hz), 8.86 (1H, d, J = 5.2 Hz), 9.27 (1H, d, J = 1.2 Hz) (CDCI3)	493
YA1576	1.45 (3H, t, J = 7.0 Hz), 3.08 (1H, dd, J = 12.5, 10.6 Hz), 3.22 (3H, m), 3.58 (3H, s), 3.62 (2H, m), 4.05 (1H, m), 4.08 (2H,q, J = 7.0 Hz), 6.98 (2H, d, J = 8.0 Hz), 7.32 (1H, s), 7.49 (2H, d, J = 8.0 Hz), 7.52 (2H, d, J = 8.0 Hz), 7.58 (2H, d, J = 8.0 Hz), 8.17 (1H, d, J = 5.3 Hz), 8.86 (1H, d, J = 5.3 Hz), 9.27 (1H, s), (CDCI3)	469
YA1577	1.83 (4H, m), 1.99 (1H, m), 2.21 (1H, m), 2.61 (4H, m), 2.87 (1H, m), 3.03 (1H, dd, J = 12.0, 10.0 Hz), 3.20 (4H, m), 3.33 (1H, m), 3.42 (1H, m), 3.49 (1H, m), 3.56 (3H, s), 3.61 (2H, m), 3.90 (1H, dd, J = 10.0, 2.0 Hz), 6.55 (2H, d, J = 8.8 Hz), 7.29 (2H, d, J = 8.8 Hz), 7.30 (1H, s), 8.16 (1H, d, J = 5.2 Hz), 8.85 (1H, d, J = 5.2 Hz), 9.26 (1H, s) (CDCI3)	487
YA1578	3.09 (1H, dd, J = 12.4, 10.8 Hz), 3.20 (3H, m), 3.58 (3H, s), 3.64 (2H, m), 3.82 (3H, s), 3.86 (3H, s), 4.05 (1H, dd, J = 10.4, 2.8 Hz), 6.58 (2H, m), 7.24 (2H, m), 7.32 (1H, s), 7.47 (2H, d, J = 8.4 Hz), 7.53 (2H, d, J = 8.4 Hz), 8.17 (1H, dd, J = 5.2, 1.2 Hz), 8.86 (1H, d, J = 5.2 Hz), 9.27 (1H, d, J = 1.2 Hz) (CDCI3)	485

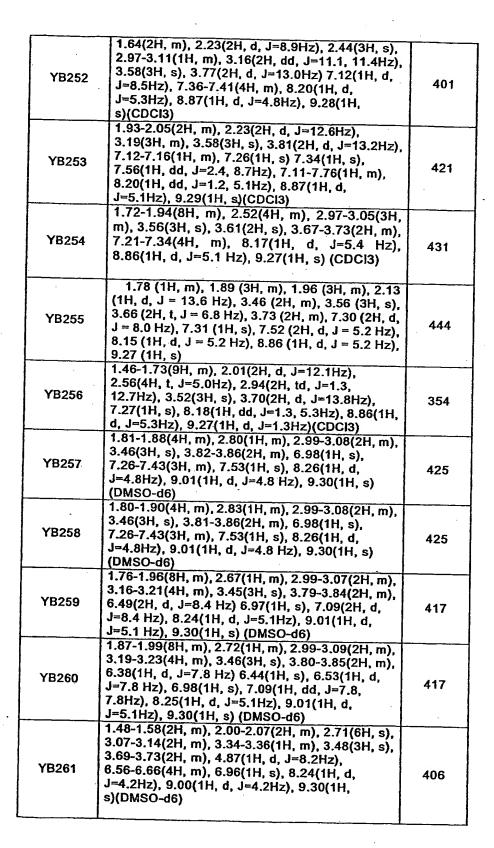
YA1579	3.08 (1H, dd, J = 12.5, 10.6 Hz), 3.23 (3H, m), 3.59 (3H, s), 3.66 (2H, m), 3.93 (3H, s), 3.96 (3H, s), 4.07 (1H, dd, J = 10.3, 2.2 Hz), 6.95 (1H, d, J = 8.3 Hz), 7.11 (1H, d, J = 2.0 Hz), 7.16 (1H, dd, J = 8.3, 2.0 Hz), 7.33 (1H, s), 7.52 (1H, d, J = 8.1 Hz), 7.59 (1H, d, J = 8.1 Hz), 8.17 (1H, dd, J = 5.3, 1.2 Hz), 8.85 (1H, d, J = 5.3 Hz), 9.27 (1H, d, J = 1.2 Hz) (CDCI3)	485
YA1580	3.07 (1H, dd, J = 12.4, 10.4 Hz), 3.23 (3H, m), 3.59 (3H, s), 3.65 (2H, m), 4.08 (1H, dd, J = 10.4, 2.0 Hz), 7.32 (1H, s), 7.41 (2H, d, J = 8.4 Hz), 7.52 (2H, d, J = 8.4 Hz), 7.53 (2H, d, J = 8.4 Hz), 7.58 (2H, d, J = 8.4 Hz), 8.16 (1H, d, J = 4.8 Hz), 8.86 (1H, d, J = 4.8 Hz), 9.27 (1H, s) (CDCI3)	459
YA1581	3.09 (1H, dd, J = 12.2, 11.0 Hz), 3.24 (3H, m), 3.59 (3H, s), 3.66 (2H, m), 4.10 (1H, dd, J = 10.4, 2.4 Hz), 7.29 (2H, m), 7.33 (1H, s), 7.44 (2H, d, J = 8.0 Hz), 7.52 (3H, m), 8.18 (1H, dd, J = 5.3, 1.0 Hz), 8.87 (1H, d, J = 5.3 Hz), 9.27 (1H, d, J = 1.0 Hz) (CDCI3)	493
YA1582	3.06 (1H, dd, J = 12.4, 10.4 Hz), 3.25 (3H, m), 3.58 (3H, s), 3.65 (2H, m), 4.09 (1H, dd, J = 10.0, 2.0 Hz), 7.33 (1H, s), 7.42 (1H, dd, J = 8.0, 2.0 Hz), 7.56 (5H, m), 7.68 (1H, d, J = 2.0 Hz), 8.16 (1H, dd, J = 5.2, 1.2 Hz), 8.85 (1H, d, J = 5.2 Hz), 9.27 (1H, d, J = 1.2 Hz) (CDCl3)	493
YA1583	3.06 (1H, dd, J = 12.3, 10.8 Hz), 3.23 (3H, m), 3.59 (3H, s), 3.65 (2H, m), 4.13 (1H, dd, J = 10.2, 2.2 Hz), 7.33 (1H, s), 8.14 (1H, d, J = 5.3 Hz), 8.15 (2H, d, J = 8.4 Hz), 8.78 (1H, s), 8.86 (1H, d, J = 5.3 Hz), 9.27 (1H, s) (CDCl3)	417
YA1584	1.37(6H, d, J= 6.0 Hz), 3.07(1H, dd, J=12.6, 10.8 Hz), 3.20-3.26(3H, m), 3.58(3H, s), 3.65-3.68(2H, m), 4.07(1H, m), 4.59(1H, m), 6.98(2H, d, J= 8.7 Hz), 7.48(1H, s), 7.50-7.61(6H, m), 8.17(1H, d, J=4.8 Hz), 8.86(1H, d, J=5.1 Hz), 9.26(1H, d, J=1.2 Hz) (CDCI3)	483
YA1585	0.99(3H, t, J= 7.5 Hz), 1.47-1.82(4H, m), 3.07(1H, dd, J=12.3, 10.5 Hz), 3.22-3.27(3H, m), 3.58(3H, s), 3.62-3.65(2H, m), 4.03(2H, t, J= 6.3 Hz), 4.04(1H, m), 6.98(2H, d, J= 8.7 Hz), 7.48(1H, s), 7.50-7.59(6H, m), 8.17(1H, dd, J=5.1, 1.2 Hz), 8.86(1H, d, J=5.1 Hz), 9.26(1H, d, J=1.2 Hz) (CDCI3)	497
YA1586	1.28(1H, br.s), 2.51(3H, s), 3.07(1H, dd, J=10.8, 12.6Hz), 3.21-3.28(3H, m), 3.58(3H, s), 3.64(2H, m), 4.08(1H, dd, J=2.5, 19.5Hz), 7.34(2H, d, J=7.8Hz), 7.45-7.67(7H, m), 8.17(1H, d, J=5.4Hz), 8.86(1H, d, J=5.1Hz), 9.27(1H, d, J=1.2Hz)(CDCI3)	470



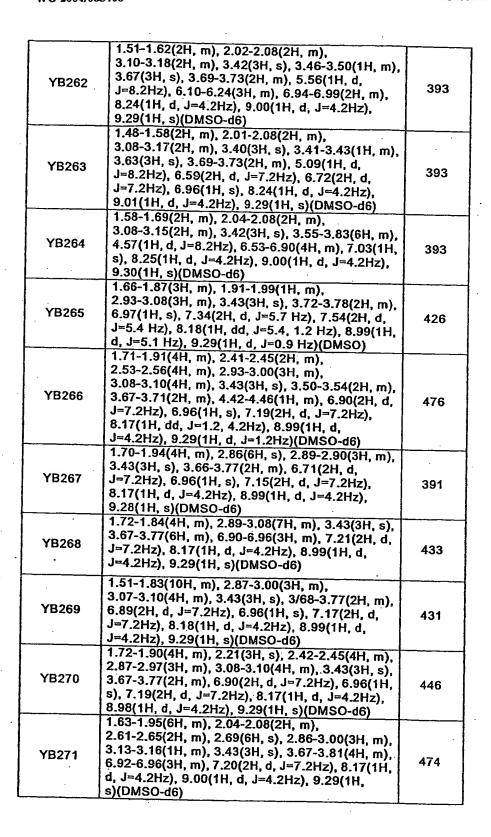
#### WO 2004/085408



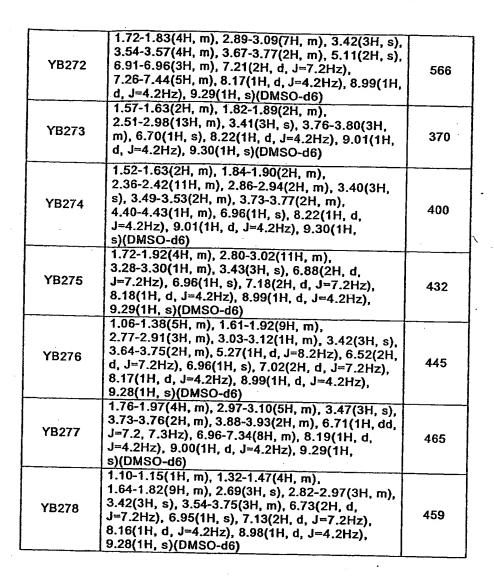








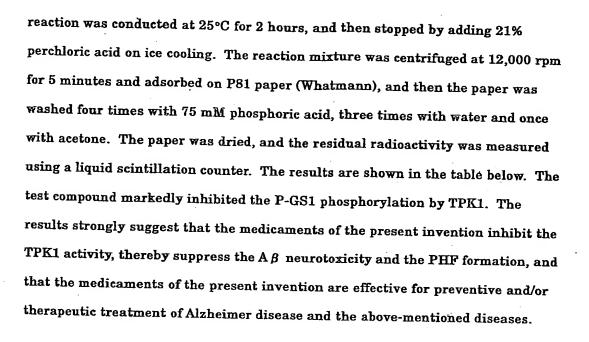




Test Example: Inhibitory activity of the medicament of the present invention against P-GS1 phosphorylation by bovine cerebral TPK1

A mixture containing 100 mM MES-sodium hydroxide (pH 6.5), 1 mM magnesium acetate, 0.5 mM EGTA, 5 mM  $\beta$ -mercaptoethanol, 0.02% Tween 20, 10% glycerol, 12  $\mu$  g/ml P-GS1, 41.7  $\mu$  M [ $\gamma$ -32P] ATP (68 kBq/ml), bovine cerebral TPK1 and a compound shown in Table (a final mixture contained 1.7% DMSO deriving from a solution of a test compound prepared in the presence of 10% DMSO) was used as a reaction system. The phosphorylation was started by adding ATP, and the









#### WO 2004/085408

Table 6

Compound No.	IC50
XA361	0.018 μ M
XB80	0.23 μ Μ
ŸA0864	0.216 μ M
YB257	0.014 μ M

### Formulation Example

### (1) Tablets

The ingredients below were mixed by an ordinary method and compressed by using a conventional apparatus.

Compound of Example 1	30 mg
Crystalline cellulose	60 mg
Corn starch	100 mg
Lactose	200 mg
Magnesium stearate	4 mg

### (2) Soft capsules

The ingredients below were mixed by an ordinary method and filled in soft capsules.

Compound of Example 1	30  mg
Olive oil	300 mg
Lecithin	20 mg

## Industrial Applicability

The compounds of the present invention have TPK1 inhibitory activity and are useful as an active ingredient of a medicament for preventive and/or therapeutic treatment of diseases caused by abnormal advance of TPK1 such as neurodegenerative diseases (e.g. Alzheimer disease) and the above-mentioned diseases.



### CLAIMS

 A pyrimidone derivative represented by formula (I) or a salt thereof, or a solvate thereof or a hydrate thereof:

$$(X)_{m}$$

$$(X)_{m}$$

$$(X)_{n}$$

$$(X)_$$

wherein Q represents CH or nitrogen atom;

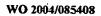
R represents a C<sub>1</sub>-C<sub>12</sub> alkyl group which may be substituted; the ring of:

represents piperazine ring or piperidine ring;

each X independently represents

$$X^1 - X^2 -$$

wherein X¹ represents an oxo group; a C¹-C³ alkyl group which may be substituted; a C³-C³ cycloalkyl group which may be substituted; an optionally partially hydrogenated C⁵-C¹⁰ aryl ring which may be substituted; an indan ring which may be substituted; an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total; an aralkyloxy group; a group represented by -N(R³)(R³) wherein R³ and R³ are the same or different and each is hydrogen, a C¹-C⁴ alkyl group which may be substituted, an aralkyl group which may be substituted, an





aryl group which may be substituted, C1-C8 alkylcarbonyl group which may be substituted,

C3-C3 cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted,
C6-C10 arylcarbonyl group which may be substituted,
C1-C3 alkysulfonyl group which may be substituted,
C3-C3 cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted,
C6-C10 arylsulfonyl group which may be substituted,
C1-C3 alkyloxycarbonyl group which may be substituted,
C3-C3 cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted,
C6-C10 aryloxycarbonyl group which may be substituted,
C6-C10 aryloxycarbonyl group which may be substituted, aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted,
N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,
N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,
N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,
N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,
C3-C8 cycloalkylaminocarbonyl group which may be substituted,
N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,
N-C3-C8 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,
N-C3-C8 cycloalkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,
aralkylaminocarbonyl group which may be substituted,
N,N'-diaralkylaminocarbonyl group which may be substituted,
N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

Ce-C10 arylaminocarbonyl group which may be substituted,

N,N'-C6-C10 diarylaminocarbonyl group which may be substituted,



or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total; or Ra and Rb together with the adjacent nitrogen atom form a 4 to 7 membered heterocyclic ring which may further contain 1 to 4 groups selected from an oxygen atom, a sulfur atom, N-Rc (wherein Rc represents a hydrogen atom, a C1-C4 alkyl group which may be substituted, an aralkyl group which may be substituted,  $C_3$ - $C_8$  cycloalkyl group which may be substituted or an aryl group which may be substituted, C<sub>1</sub>-C<sub>8</sub> alkylcarbonyl group which may be substituted, C3-C8 cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted, C6-C10 arylcarbonyl group which may be substituted, C1-C8 alkysulfonyl group which may be substituted, C<sub>3</sub>-C<sub>8</sub> cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted, C6-C10 arylsulfonyl group which may be substituted, C<sub>1</sub>-C<sub>8</sub> alkyloxycarbonyl group which may be substituted, C<sub>8</sub>-C<sub>8</sub> cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted,  $C_6$ - $C_{10}$  aryloxycarbonyl group which may be substituted, aminocarbonyl, N-C1-C8 alkylaminocarbonyl group which may be substituted, N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,
N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,
N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,
C3-C8 cycloalkylaminocarbonyl group which may be substituted,
N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted, N-C<sub>5</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted, N,N'-diaralkylaminocarbonyl group which may be substituted, N-aralkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, N,N'-C<sub>6</sub>-C<sub>10</sub> diarylaminocarbonyl group which may be substituted, or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and having 5 to 10 ring-constituting atoms in total),

a carbonyl group, a sulfinyl group or a sulfonyl group in the ring, and said 4 to 7 membered heterocyclic ring may optionally be fused with an aryl group which may be substituted;

X² represents a bond, a carbonyl group, a sulfinyl group, a sulfonyl group, an oxygen atom, a sulfur atom, a C₁-C₄ alkylene group which may be substituted or N-Rd (Rd represents a hydrogen atom, a C₁-C₄ alkyl group which may be substituted, an aralkyl group which may be substituted, C₃-C₃ cycloalkyl group which may be substituted or an aryl group which may be substituted,
C₁-C₃ alkylcarbonyl group which may be substituted,
C₃-C₃ cycloalkylcarbonyl group which may be substituted,
aralkycarbonyl group which may be substituted,
C₁-C₃ alkysulfonyl group which may be substituted,
C₁-C₃ alkysulfonyl group which may be substituted,
C₃-C₃ cycloalkylsulfonyl group which may be substituted,
aralkysulfonyl group which may be substituted,
C₃-C₃ cycloalkylsulfonyl group which may be substituted,

WO 2004/085408 PCT/JP2004/004320

C<sub>1</sub>-C<sub>8</sub> alkyloxycarbonyl group which may be substituted,
C<sub>2</sub>-C<sub>8</sub> cycloalkyloxycarbonyl group which may be substituted,
aralkyoxycarbonyl group which may be substituted,
C<sub>6</sub>-C<sub>10</sub> aryloxycarbonyl group which may be substituted,
aminocarbonyl,

N-C1-C8 alkylaminocarbonyl group which may be substituted, N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted, N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted, N-C<sub>1</sub>-C<sub>8</sub> alkyl-N'-aralkylaminocarbonyl group which may be substituted, N-C<sub>1</sub>-C<sub>8</sub> alkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, C3-C8 cycloalkylaminocarbonyl group which may be substituted, N,N'-C<sub>8</sub>-C<sub>8</sub> dicycloalkylaminoycarbonyl group which may be substituted, N-C3-C8 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted, N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted, N,N'-diaralkylaminocarbonyl group which may be substituted, N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted, C6-C10 arylaminocarbonyl group which may be substituted, N,N'-C6-C10 diarylaminocarbonyl group which may be substituted, or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and

m represents an integer of 1 to 3;

having 5 to 10 ring-constituting atoms in total);

each Y independently represents a halogen atom, a hydroxy group, a cyano group, Y1-Y3- wherein Y1 represents a C1-C8 alkyl group which may be substituted; a C3-C8 cycloalkyl group which may be substituted or a C6-C10 aryl ring which may be substituted; Y3 represents a carbonyl group, a sulfinyl group, a sulfonyl group, an oxygen atom, a sulfur atom, a C1-C4 alkylene group which may be substituted or





WO 2004/085408

N-Re (Re represents a hydrogen atom, a C1-C4 alkyl group which may be substituted, an aralkyl group which may be substituted, C3-C8 cycloalkyl group which may be substituted or an aryl group which may be substituted, C1-C8 alkylcarbonyl group which may be substituted, Cs-Cs cycloalkylcarbonyl group which may be substituted, aralkycarbonyl group which may be substituted, C6-C10 arylcarbonyl group which may be substituted, C1-C8 alkysulfonyl group which may be substituted, C3-C8 cycloalkylsulfonyl group which may be substituted, aralkysulfonyl group which may be substituted, C6-C10 arylsulfonyl group which may be substituted, C1-C8 alkyloxycarbonyl group which may be substituted, C3-C8 cycloalkyloxycarbonyl group which may be substituted, aralkyoxycarbonyl group which may be substituted, C6-C10 aryloxycarbonyl group which may be substituted, aminocarbonyl, N-C1-C8 alkylaminocarbonyl group which may be substituted,

N, N'-C1-C8 dialkylaminocarbonyl group which may be substituted.

N-C1-C8 alkyl-N'-C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C1-C8 alkyl-N'-C6-C10 arylaminocarbonyl group which may be substituted,

C3-C8 cycloalkylaminocarbonyl group which may be substituted,

N,N'-C3-C8 dicycloalkylaminoycarbonyl group which may be substituted,

N-C3-C8 cycloalkyl-N'-aralkylaminocarbonyl group which may be substituted,

N-C<sub>3</sub>-C<sub>8</sub> cycloalkyl-N'-C<sub>6</sub>-C<sub>10</sub> arylaminocarbonyl group which may be substituted, aralkylaminocarbonyl group which may be substituted,

N,N'-diaralkylaminocarbonyl group which may be substituted,

N-aralkyl- N'-C6-C10 arylaminocarbonyl group which may be substituted,

WO 2004/085408 PCT/JP2004/004320

C6-C10 arylaminocarbonyl group which may be substituted,

N,N'-C6-C10 diarylaminocarbonyl group which may be substituted,

or an optionally substituted heterocyclic ring having 1 to 4 hetero atoms selected
from the group consisting of oxygen atom, sulfur atom, and nitrogen atom, and
having 5 to 10 ring-constituting atoms in total),

n represents an integer of 0 to 8;

when X and Y or two Y groups are attached on the same carbon atom, they may
combine to each other to form a C2-C6 alkylene group;
and when m is 1, n is 0, and X is X1-CO-,

- (1) X does not bind to 3-position of unsubstituted 1-piperazinyl group or does not bind to 3-position of a 4-alkyl-1-piperazinyl group; or
- (2) X does not bind to 3-position or 4-position of non-substituted 1-piperidinyl group.
- 2. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 1 having the following formula(II)

$$(X)_{p}$$

$$(X)_{q}$$

$$(Y)_{r}$$

$$(X)_{q}$$

$$(Y)_{r}$$

$$(X)_{q}$$

$$(Y)_{r}$$

wherein Q, R, X and Y are the same as those defined in claim 1; p is 0 or 1; q is 0 or 1; r is an integer of 0 to 6; p+q is 1 or 2; and Z represents N or CZ<sup>1</sup> wherein Z<sup>1</sup> represents hydrogen atom or Y.

3. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 2, wherein R is a C<sub>1</sub>-C<sub>3</sub> alkyl group which





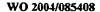
may be substituted by a C3-C8 cycloalkyl group.

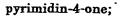
- 4. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 3, wherein R is methyl group or ethyl group; Y is in 3-, 4- or 5-position of the piperazine ring or the piperidine ring; p+q is 1; and r is an integer of 0 to 3.
- 5. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 4, wherein X is a C<sub>1</sub>-C<sub>8</sub> alkyl group which may be substituted or a C<sub>8</sub>-C<sub>10</sub> aryl ring which may be substituted; Y is a C<sub>1</sub>-C<sub>6</sub> alkyl group which may be substituted; p is 1; q is 0; r is an integer of 0 to 3; and Z is N or CH.
- 6. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 5, wherein X is a benzene ring which may be substituted, a benzyl group which may be substituted; Y is a methyl group which may be substituted; Z is N and r is 0 or 1.
- 7. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 4, wherein X is a benzene ring which may be substituted, a benzyl group which may be substituted, a benzyl group which may be substituted; Y is a methyl group which may be substituted; Z is N and p is 0.
- 8. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 4, wherein X is a C<sub>1</sub>-C<sub>8</sub> alkyl group substituted by a benzene ring which may be substituted or a benzene ring which may be substituted; Y is a hydroxy group, a cyano group, or Y¹-CO- wherein Y¹ is a C¹-C³ alkyl group; Z is CH or C-Y and r is 0 or 1.
- 9. The pyrimidone derivative or the salt thereof, or the solvate thereof or the hydrate thereof according to claim 8, wherein X is a benzyl group which may be substituted or a benzene ring which may be substituted; Y is a hydroxy group, a cyano group, or an acetyl group; Z is CH or C-Y and r is 0 or 1.





2-(3-(4-Fluoro-3-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-





- 2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Chloro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methylphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Fluoro-6-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(5-Bromo-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Bromo-4-fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Chloro-6-fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,4-Difluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Difluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dichlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3,4-Dimethoxyphenyl) piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-

4-one;

- 2-(3-(2,5-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dimethoxyphenyl)piperasin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,4-Difluoro-6-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;(1034)
- 2-(3-(5-Cyano-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Cyano-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(1-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2,3-Dihydrobenzofuran-7-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- $\hbox{2-(3-(Benzo furan-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} \\ H-pyrimidin-4-one;$
- (S)-2-(3-(Benzofuran-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-4-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-5-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Phenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one; 2-(3-(4-(4-Fluorophenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-



# pyrimidin-4-one;

pyrimidin-4-one;

WO 2004/085408

- $\hbox{2-(3-(4-(4-Methoxyphenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3} H-10-(4-Methoxyphenyl)$
- 2-(3-(4-(2-Methoxyphenyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Morpholin-4-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(4-Methylpiperazin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Phenylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzoylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-(1,2-Benzisothiazol-3-yl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Methyl-3-phenylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)-4-methylpiperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(4\hbox{-}Acetyl\hbox{-}3\hbox{-}(4\hbox{-}fluoro\hbox{-}2\hbox{-}methoxyphenyl) piperazin-1-yl)\hbox{-}3\hbox{-}methyl\hbox{-}6\hbox{-}(4\hbox{-}pyridyl)\hbox{-}3H-pyrimidin-4\hbox{-}one;}$
- 2-(4-Benzyl-3-(4-fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-Benzyl-3-(ethoxycarbonyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(4-methyl-3-(1-naphthyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-

one;

- 2-(5,5-Dimethyl-3-(2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-Phenylpiperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Fluorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Chlorophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Bromophenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Methoxyphenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3-Methoxyphenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Methoxyphenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(4-((Pyrrolidin-1-yl)methyl)phenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-
- pyrimidin-4-one;
- (S)-2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-
- 3H-pyrimidin-4-one;
- (R)-2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperidin-1-yl)-3-methyl-6-(4-pyridyl)-
- 3H-pyrimidin-4-one;
- 2-(3-Hydroxy-3-phenylpiperidin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-Phenylpiperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(3-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Fluorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Chlorophenyl) piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-<math>4-yl
- one;
- $\hbox{2-(3-(3-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3} H-pyrimidin-4-pyrimidyl-3 H-pyrimidyl-4-pyrimidyl-3 H-pyrimidyl-4-pyrimidyl-3 H-pyrimidyl-4-pyrimidyl-6 H-pyrimidyl-6 H-pyrimidyl$
- one;
- $\hbox{2-(3-(2-Chlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3$$H$-pyrimidin-4-pyrimidyl-3$$H$-pyrimidyl-3$$H$-pyrimidyl-3$$H$-pyrimidyl-4$$H$-pyrimidyl-3$$H$-pyrimidyl-4$$H$-pyrimidyl-3$$H$-pyrimidyl-4$$H$-pyrimidyl-$

one;

- 2-(3-(4-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(3-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(2-Bromophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(4-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(3-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(2-Cyanophenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one; 2-(3-(4-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(3-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2-Ethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(6-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(5-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (S)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- (R)-2-(3-(4-Fluoro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-Chloro-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- $\hbox{2-(3-(5-Bromo-2-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3} \\ H-10-2-methyl-6-(4-pyrimidyl)-3 \\ H-10-2-methyl-6-(4-pyrimidyl$





### pyrimidin-4-one;

- 2-(3-(2,6-Dichlorophenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-
- 2-(3-(2,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyridyl)-3H-pyrimidin-4-one;
- 2-(3-(3,4-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,5-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2,6-Dimethoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one;
- 2-(3-(2,4-Difluoro-6-methoxyphenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(1\hbox{-Naphthyl}) piperazin-1\hbox{-}yl)\hbox{-}3\hbox{-methyl-}6\hbox{-}(4\hbox{-pyrimidyl})\hbox{-}3H\hbox{-pyrimidin-}4\hbox{-}one;$
- $2\hbox{-}(3\hbox{-}(2\hbox{-Naphthyl}) piperazin-1\hbox{--yl})-3\hbox{--methyl-}6\hbox{-}(4\hbox{--pyrimidyl})-3H\hbox{--pyrimidin-}4\hbox{--one};$
- 2-(3-(2,3-Dihydrobenzofuran-7-yl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(Benzofuran-2-yl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl-methyl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(4-(Pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-4-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- 2-(3-(2-methoxy-5-(pyrrolidin-1-yl)phenyl)piperazin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;
- $2\hbox{-}(3\hbox{-}(4\hbox{-}(Phenyl)phenyl)piperazin-1-yl)-3-methyl-6\hbox{-}(4\hbox{-}pyrimidyl)-3$H-pyrimidin-4-pyrimidyl-3$H-pyrimidyl-4-pyr$

pyrimidin-4-one;

(R)-2-(3-(Benzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;

2-(3-(6-Fluorobenzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3H-pyrimidin-4-one;

2-(4-(6-Fluorobenzoisoxazol-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one;

2-(4-(5-Methylbenzofuran-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one; and

2-(4-(6-Fluorobenzothiophene-3-yl)piperidin-1-yl)-3-methyl-6-(4-pyrimidyl)-3*H*-pyrimidin-4-one

or a salt thereof, or a solvate thereof or a hydrate thereof.

- 11. A medicament comprising as an active ingredient a substance selected from the group consisting of the pyrimidone derivative represented by formula (I) and a salt thereof, and a solvate thereof and a hydrate thereof according to claim 1.
- 12. A tau protein kinase 1 inhibitor selected from the group consisting of the pyrimidone derivative represented by formula (I) and a salt thereof, and a solvate thereof and a hydrate thereof according to claim 1.
- 13. The medicament according to claim 11 which is used for preventive and/or therapeutic treatment of a disease caused by tau protein kinase 1 hyperactivity.
- 14. The medicament according to claim 11 which is used for preventive and/or therapeutic treatment of a neurodegenerative disease.
- 15. The medicament according to claim 14, wherein the neurodegenerative disease is selected from the group consisting of Alzheimer disease, ischemic cerebrovascular accidents, Down syndrome, cerebral bleeding due to cerebral amyloid angiopathy, progressive supranuclear palsy, subacute sclerosing panencephalitic parkinsonism, postencephalitic parkinsonism, pugilistic

encephalitis, Guam parkinsonism-dementia complex, Lewy body disease, Pick's disease, corticobasal degeneration, frontotemporal dementia, vascular dementia, traumatic injuries, brain and spinal cord trauma, peripheral neuropathies, retinopathies, and glaucoma.

16. The medicament according to claim 11, wherein the disease is selected from the group consisting of non-insulin dependent diabetes, obesity, manic depressive illness, schizophrenia, alopecia, breast cancer, non-small cell lung carcinoma, thyroid cancer, T or B-cell leukemia, and a virus-induced tumor.



### INTERNATIONAL SEARCH REPORT

mational Application No PCT/JP2004/004320

A CLASSIFICATION OF SUBJECT MATTER

IPC 7 C07D239/47 C07D401/14 CO7D409/14 C07D405/14 CO7D413/14 CO7D417/14 CO7D403/14 A61K31/513 A61K31/5377 A61P25/28

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) CO7D A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, BEILSTEIN Data, WPI Data, PAJ, CHEM ABS Data

C. DOCUM	ENTS CONSIDERED TO BE RELEVANT		
Category •	Citation of document, with indication, where appropriate, of	Relevant to claim No.	
X	EP 1 136 482 A (SANOFI SYNTHE MITSUBISHI TOKYO PHARMACEUTIC 26 September 2001 (2001-09-26 cited in the application paragraphs '0002!, '0006!; e tables 1,2,4,8-12	1-16	
P,A	WO 03/027080 A (SANOFI SYNTHE SHINSUKE (JP); SHODA AYA (JP) KEIICH) 3 April 2003 (2003-04 page 104, line 9 - line 25; c 1,2,10-25; examples C401-C499,C601-C651,C751-C768	1-16	
	<del></del>	-/·	
		,	
			'
	·		
X Funi	her documents are listed in the continuation of box C.	X Palent family member	rs are listed in annex.
*Special categories of cited documents:  'A' document defining the general state of the art which is not considered to be of particular relevance  'E' earlier document but published on or after the international filling date  'L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  'O' document reterring to an oral disclosure, use, exhibition or other means  'P' document published prior to the international filling date but later than the priority date claimed  Date of the actual completion of the international search		or priority date and not in cited to understand the prinvention  "X" document of particular relections involve an inventive step  "Y" document of particular relections of particular relections involve an inventive step  "Y" document of particular relections of particular relections in the particular relec	rel or cannot be considered to when the document is taken alone avance; the claimed invention involve an inventive step when the tith one or more other such docubeing obvious to a person skilled same patent family
Name and r	mailing address of the ISA	Authorized officer	
	European Patent Office, P.B. 5818 Patentiaan 2 NL - 2260 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Hanisch, I	·





### INTERNATIONAL SEARCH REPORT

mational Application No PCT/JP2004/004320

Category •	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
ALLEGUTY -	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
P,A	WO 03/037888 A (SANOFI SYNTHELABO; HIKI SHINSUKE (JP); SHODA AYA (JP); ARITOMO KEIICH) 8 May 2003 (2003-05-08) page 9, line 21 - page 10, line 10; claims 1,2,11-13,15-25; examples C001-C099,C201-C251,C351-C368,C388,D001-D0 25	1-16	
	**************************************		
,			
1			
	•		
		<u>.</u>	
		1	
	·	1	

# INTERNATIONAL SEARCH REPORT

information on patent family members

ational Application No

		r		207/2	application No
Patent document cited in search report		Publication date	T	Patent tamily	2004/004320
EP;1135482	I	26-09-2001	┸	member(s)	Publication date
). 1	A	20-09-2001	EP AU CA CN WO EP JP US	1136482 A1 8148701 A 2401241 A1 1419552 T 0170729 A1 1276738 A1 2003528095 T 2003187004 A1	26-09-2001 03-10-2001 27-09-2001 21-05-2003 27-09-2001 22-01-2003 24-09-2003
WO 03027080	A	03-04-2003	CA CA EP EP WO	2460121 A1 2460177 A1 1427709 A1 1427720 A1 03027080 A1 03037888 A1	02-10-2003 08-05-2003 03-04-2003 16-06-2004 16-06-2004 03-04-2003
WO 03037888	A	08-05-2003	CA CA EP EP WO	2460121 A1 2460177 A1 1427709 A1 1427720 A1 03027080 A1 03037888 A1	08-05-2003 08-05-2003 03-04-2003 16-06-2004 16-06-2004 03-04-2003 08-05-2003

# This Page is Inserted by IFW Indexing and Scanning Operations and is not part of the Official Record

### **BEST AVAILABLE IMAGES**

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

☐ BLACK BORDERS
☐ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
FADED TEXT OR DRAWING
☐ BLURRED OR ILLEGIBLE TEXT OR DRAWING
☐ SKEWED/SLANTED IMAGES
☐ COLOR OR BLACK AND WHITE PHOTOGRAPHS
☐ GRAY SCALE DOCUMENTS
☐ LINES OR MARKS ON ORIGINAL DOCUMENT
☐ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY
OTHER.

# IMAGES ARE BEST AVAILABLE COPY.

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.